

PCT/US2005/027239

moving from start (N-terminus or 5') to end (C-terminus or 3'), such that for an alignment that extends to  $p$  monomers (where  $p > x$ ) there are  $p-x+1$  such windows, each window has at least  $x \cdot y$  identical aligned monomers, where:  $x$  is selected from 20, 25, 30, 35, 40, 45, 50, 60, 70, 80, 90, 100, 150, 200;  $y$  is selected from 0.50, 0.60, 0.70, 0.75, 0.80, 0.85, 0.90, 0.91, 0.92, 0.93, 0.94, 0.95, 0.96, 0.97, 0.98, 0.99; and if  $x \cdot y$  is not an integer then it is rounded up to the nearest integer. The preferred pairwise alignment algorithm is the Needleman-Wunsch global alignment algorithm [Needleman & Wunsch (1970) *J. Mol. Biol.* 48, 443-453], using default parameters (e.g., with Gap opening penalty = 10.0, and with Gap extension penalty = 0.5, using the EBLOSUM62 scoring matrix). This algorithm is conveniently implemented in the *needle* tool in the EMBOSS package [Rice *et al.* (2000) *Trends Genet.* 16:276-277].

The nucleic acids and polypeptides of the invention may additionally have further sequences to the N-terminus/5' and/or C-terminus/3' of these sequences (a) to (d).

All of the Gram positive bacterial sequences referenced herein are publicly available through PubMed on GenBank.

#### Streptococcus pneumoniae Adhesin Island Sequences

As discussed above, a *S. pneumoniae* AI sequence is present in the TIGR4 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences are set forth below.

SrtD (Sp0468) is a sortase. An example of an amino acid sequence of SrtD is set forth in SEQ ID NO: 80.

#### **SEQ ID NO: 80**

MSRTKLRALLGYLLMLVACLIPIYCFGQMVLQSLGQVKGHATFVKSM TTEMYQEQQNHSLAYNQRLASQNRIVDP  
FLAEGYEVNYQVSDPDPAVYGYLSIPSLEIMEPVYLGADYHHLGMGLAHVDGTPLPLDGTGIRSVIAGHRAEPSH  
VFFRHLQDLKVGDALYYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAFGLFVLWKLARLLRGK

SrtC (Sp0467) is a sortase. An example of an amino acid sequence of SrtC is set forth in SEQ ID NO: 81.

#### **SEQ ID NO: 81**

MSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAFNATLKPSEILDPFTEQEKKKGVSEYANMLKVHERIG  
YVEIPAIQEIIPMYVGTSEDILQKGAGLLEGASLPVGGENTHTVITAHRGLPTAELFSQLDKMKKGDIFYLHVLD  
QVLAYQVDQIVTVEPNDFEPVLIQHGEDYATLLTCTPYMINSHRLLVRGKRIPYTAPIAERNRAVRERGFQFWLWL  
LLGAMAVILLLLYRVYRNRRIVKGLEKQLEGRHVKD

SrtB (SP0466) is a sortase. An example of an amino acid sequence of SrtB is set forth in SEQ ID NO: 82.

#### **SEQ ID NO: 82**

MAVMAYPLVSRLYRVESNQIADFDKEKATLDEADIDERMKLAQAFNDLNNVSGDPWSEEMKKKGRAEYARM  
LEIHERMGHVEIPVIDVDLPVYAGTAEVLLQQGAGHLEGTSLPIGGNSTHAVITAHTGLPTAKMFTDLTKLVGD  
KFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINTHRLLVRGHRIPYVAEVEEEFIAANK  
LSHLYRYLFYVAVGLIVILLWIIRRLRKKKKQPEKALKALKAAARKEVKVEDGQQ

Sp0465 is a hypothetical protein. An example of an amino acid sequence of Sp0465 is set forth in SEQ ID NO: 83.

**SEQ ID NO: 83**

MFLPFELSASLYLQTHHFIAFPNRQSYLLRETRKSHFFLIHHPF

RrgC (SP0464) is a cell wall surface anchor family protein. RrgC contains a sortase substrate motif VPXTG (SEQ ID NO: 137), shown in *italics* in SEQ ID NO: 84.

**SEQ ID NO: 84**

MISRIFFVMALCFSLVWGAHAVQAQEDHTLVLQLENYQEVVSQLP SRDGHRLQVWKLDDSYSDDRVQIVRDLHS  
WDENKLSSFKKT SFEMTFLENQIEVSHIPNGLYYVRSIIQTDAVSYPAEFLFEMTDQTVEPLVIVAKKTDMTTK  
VKLIKVDQDHNRLLEGVGFKLVS VARDVSEKEVPLIGEYRYSSSGQVGR TLYTDKNGEIFVTNLPLGN YRFKEVEP  
LAGYAVTTLD TDVQLVDHQLVTITV VNNQKLPRGNVDFMKVDGR TNTSLQGAMFKVMKEESGHYTPVLQNGKEVVV  
TSGKDGRFRVEGLE YGTYLWELQAPTGYVQLTSPVSFTIGKDTRKELVTVVKNKRPRIDVPDTGEETLYILML  
VAILLFGSGYYLT KKPNN

RrgB (Sp0463) is a cell wall surface anchor protein. RrgB contains a sortase substrate motif IPXTG (SEQ ID NO: 133), shown in *italics* in SEQ ID NO: 85.

**SEQ ID NO: 85**

MKSINKFLTMLAALLLTASSLFS AATVFAAGTTTT SVTVHKLLATDGDMDKIANELETGNYAGNKVGVL PANAKE  
IAGVMFVWTNTNNEIIDENGQTLGVNIDPQTFKL SGAMPATAMKKLTEAEGAKFNTANLPAAKYKIYEI HSLSTY  
VGEDGATLTGSKAVPIEIELPLNDVVD AHVYPKNTEAKPKIDKDFK GKANPDTPRVDKDTPVNHQVGDVVEYEIV  
TKIPALANYATANWSDRMTEGLAFNKGTVKVTVD DVALEAGDYALTEVATGFDLKLTDAGLAKVNDQNAEKT VKI  
TYSATLNDKAIVEVPESNDVTFN YGNPDHGNTPKPNKPNENGDLTLTKTWVDATGAPI PAGAEATFDLVNAQTG  
KVVQTVTLTDDKNTVTVNGLDKNTEYKFVERSI KGYSADYQEITTAGEIAVKNWKDENPKPLDPT EPKVVTYGGK  
FVKVNDKDNRLAGAEFVIANADNAGQYLARKADKVS QEEKQLVVTTKDALDRAVAAYNALTAQQQTQQE KEKVDK  
AQAAYNAAVIAANNAFEWVADKDNENVV KLVSDAQGRFEITGLLAGTYYLEETKQPAGYALLTSRQKF ETVTATSY  
SATGQGIEYTAGSGKDDATKVVNKKITIPQTGGIGTII FAVAGAAIMGIAVYAYVKNNKDEDQLA

RrgA (Sp0462) is a cell wall surface anchor protein. RrgA contains a sortase substrate motif YPXTG (SEQ ID NO: 186), indicated in *italics* in SEQ ID NO: 86.

**SEQ ID NO: 86**

MLNRETHMKKVRKIFQKAVAGLCCISQLTAFSS IVALAETPETS PAIGKVVIKETGEGGALLGD AVFELKNNTDG  
TTVSQRTEAQTGEAIFSNIKPGTYTLTEAQPPVGYK PSTKQWTVVEVEKNGRTTVQGEQVENREEALSDQY PQTGT  
YPDVQTPYQIIKVDGSEKNGQH KALNPNPYERVIPEG TLSKRIYQVNNLDDNQYGIELTVSGKT VYEQKDKSVPL  
DVVILLDNSNSMSNIRNKNARRAERAGEATRS LIDKITS DSENVALVTYASTIFDGTEFTVEKGVADKNGKRLN  
DSLFWNYDQTSFTTNTKDYSYLKLTNDKNDI VELKNKVPTEAEDHDGNRLMYQFGATFTQKALMKADEILTQ QAR  
QNSQKVI FHI TDGVPTMSYPINFN HATFAPSYQNQLNAFFSKSPNKDGILLSDFITQATS GEHTIVRGDGQSYQM  
FTDKTVYEKGAPAAFPVKPEKYSEMKAAGYAVIGDPINGGYIWLNWRESILAYPFNSNTAKITNHGDPTRWYNG  
NIAPDGYDVFTVGIGINGDPGTDEATATSEFMQSISSK PENYTNVTDTTKILEQLNRYFHTIVTEKKS IENG TITD  
PMGELIDLQLGTDGRFDPADYTLTANDGSRL ENGQAVGGPQNDGGLLKNKAVLYDTTEKRIRVTGLYLGTDEKVT  
LTYNVRLNDEFVSNKFYDTNGR TTHLPKEVEQNTVRDFPIPKIRDVRKYPEITISKEKKLGDI EFIKVNKNDKKP  
LRGAVFSLQKQHPDYPDIYGAIDQNGTYQNVRTGEDGKLTFKNLS DGKYRLFENSEPAGYKPVQNKPIVAFQIVN  
GEVRDVT SIVPQDI PAGYEFTNDKH YITNEPIPPKREYPR TGGIGMLPFYLG CMMMGGVLLYTRKHP

RlrA (Sp0461) is a transcriptional regulator. An example of an amino acid sequence for RlrA is set forth in SEQ ID NO: 87.

**SEQ ID NO: 87**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDTQQVQLIEHHS HQ  
TNYYFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISIATGYRVRQKCGLLLRVGLDLVKNQVVGPEYRIRF  
LIALLQFHFGEIYDLNDGSM DWVTHMIVQSNSQLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
LKNLFMYPI LMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQLILQHTRGKHL LSKF  
KNILGNDISNSLSFLTALTFLTRTFLEGLQNLVPYYNYEYHYGIESDKPLYHISK AIVQEWMT EQKIEGVIDQHR  
LYLFSLYLTETIFSSLPAIPIFIILNNQADVNLIKSIILRNFTDKVASVTGYNILISPPPSEEH LTEPLIIITTK  
EYLPYVKKQYPKGKHHFLTIALDLHVSQQRLIYQTI VDIRKEAFDKRVAMIAKKAHYLL

As discussed above, a *S. pneumoniae* AI sequence is present in the *S. pneumoniae* strain 670 genome. Examples of *S. pneumoniae* AI sequences are set forth below.



Orf1\_670 is a transposase. An example of an amino acid sequence of orf1\_670 is set forth in SEQ ID NO: 171.

**SEQ ID NO: 171**

MEHINHTTLLIGIKDKNITLNKAIQHDTHIEVFATLDYHPPKCKHCKGKQIKYDFQKPSKIPFIEIGGFPSLIHL  
 KKRRFQCKSCRKVTVAETTLVQKNCQISEMVRQKIAQLLLNREALTHIASKLAISTSTSTVYRKLKQFHQEDYT  
 TLPEILSWDEFYSYQKGKLAFAAQDFNTKKIMTILDNRROTIRNHFFKYSKEARKKVKVTVDMMSGSYIPLIKKL  
 FPNAKIVLDRFHIVQHMSRALNQTRINIMKQFDDKSLEYRALKYYWKFILKDSRKLSLKPFYARTFRETLTTPREC  
 LKKIFTLVPELKDYYDLYQLLLFHLQEKNTDQFWGLIQDTLPHLNRTFKTTLSTFICYKNYITNAIELPYSNAKL  
 EATNKLIKDIKRNAFGERNFENFKKRIFIALNIKKERTKFVLSRA

Orf2\_670 is a transcriptional regulator. An example of an amino acid sequence of Orf2\_670 is set forth in SEQ ID NO: 172.

**SEQ ID NO: 172**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDTQQVQLIEHHSHQ  
 TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISIATGYRVRQKCGLLLSVGLDLVKNQVVGPEYRIRF  
 LIALLQFHFGIEIYDLNDGSMWVTHMIVQNSQLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
 LKNLFMYPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKTHTIQLILQHTRGKHLLSKF  
 KNILGNDISNSLSFLTALTFLTRTFLEGLQNLVPYYNYEHYGIESDKPLYHISKAIVQEWMTQKIEGVIDQHR  
 LYLFSLYLTETIFSSLPAIPIFIILNNQADVNLIKSIILRNFTDKVASVTGYNILISPPPSEEHLEPLIIITTK  
 EYLPYVKKQYPKGKHHFLTIALDLHVSQQRLIYQTIVDIRKEAFDKRVAMIAKKAHYLL

Orf3\_670 is a cell wall surface anchor family protein. An example of an amino acid sequence of Orf3\_670 is set forth in SEQ ID NO: 173.

**SEQ ID NO: 173**

MLNRETHMKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETSIPAIGKVVIKETGEGGALLGDAVFELKNNTDG  
 TTVSQRTAEQTGEAIFSNIKPGTYTLTEAQPPVGYPSTKQWTVVEVEKNGRTTVQGEQVENREEALSDQYPQTGT  
 YPDVQTPYQIIKVDGSEKNGQHKALNPYPYERVIPEGTLISKRIYQVNNLDDNQYGIELTVSGKTTVETKEASTPL  
 DVVILLDNSNSMSNIRHNHAHRAEKAGEATRALVDKITSNPDNRVALVTYGSTIFDGSEATVEKGVADANGKILN  
 DSALWTFDRTTFTAKTYNYSFLNLTSDPDTIQTIKDRIPSDAEELNKDKLMYQFGATFTQKALMTADDILTQKAR  
 PNSKKVIFHITDGVPTMSYPINFKYTGTTQSYRTQLNNFKAKTPNSSGILLEDFVTWSADGEHKIVRGDGESYQM  
 FTKKPVTDQYGVHQILSITSMEQRAKLVSAGYRFYGTDLVLYWRDSILAYPFNSSTDWITNHGDPTTWYNGNMA  
 QDGYDVFTVGVGVNGDPGTDEATATRFMQSISSSPDNYTNVADPSQILQELNRYFYTTIVNEKKSIENTITDPMG  
 ELIDFQLGADGRFDPADYTLTANDGSSLVNNVPTGGPQNDGGLLKNKAKVFYDTTEKRIRVTGLYLGTGEKVTLT  
 NVRLNDQFVSNKFYDTNGRITLHPKEVEKNTVRDFPIPKIRDVRKYPEITIPKEKKLGEIEFIKINKNDKKPLRD  
 AVFSLQKQHPDYPDIYGAIDQNGTYQNVRTGEDGKLTFKNLSDGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEV  
 RDVTSIVPQDIPAGYEFTNDKHYITNEPIPPKREYPRGTGGIGMLPFYILIGCMGMMGGVLLYTRKHP

Orf4\_670 is a cell wall surface anchor family protein. An example of an amino acid sequence of orf4\_670 is set forth in SEQ ID NO: 174.

**SEQ ID NO: 174**

MKSINKFLTMLAALLLTASSLFSAAATVFAADNVSTAPDAVTKTLTIHKLKLLSEDDLKTWDTNGPKGYDGTQSSSLK  
 DLTGVVAEEIPNVYFELQKYNLTDGKEKENLKDDSKWTTVHGGLTTKDGLKIETSTLKGVYRIREDRTKTTYVGP  
 NGQVLTGSKAVPALVTLPLVNNNGTVIDAHVFPKNSYNKPVVDKRIADTLNNDQNGLSIGTKIPYVNTTIPSN  
 ATFATSFWSDEMTEGLTYNEDVTITLNNVAMDQADYEVTKGNNGFNLKLTAEAGLAKINGKDADQKIQTYSATLN  
 SLAVADIPESNDITYHYGNHQDHGNTPKPTKPNNGQITVTKTWSQPAPEGVKATVQLVNAKTGEKVGAPVELSE  
 NNWYTWSGLDNSIEYKVEEYNGYSAEYTVESKGLGVKNWKNPAPINPEEPRVKTYGKKFVKVDQKDTRLE  
 NAQFVVKKADS NKYIAFKSTAQQAADEKAAATAKQKLDAVAAYTNAADKQAAQALVDQAQQEYNVAYKEAKFGY  
 VEVAGKDEAMVLTSNTDGQFQISGLAAGTYKLEEIKAPEGFAKIDDFEVVGAGSWNQGEFNYLKDVQKNDATKV  
 VNKKITIPQTGGIGTIIFAVAGAAIMGIAVYAYVKNKDEDQLA

Orf5\_670 is a cell wall surface anchor family protein. An example of an amino acid sequence of orf5\_670 is set forth in SEQ ID NO: 175.

**SEQ ID NO: 175**

MTMQKMQMISRIFFVMALCFSLVWGAHAVQAQEDHTLVQLQENYQEVVSQLP SRDGHRLQVWKLDDSYSDDRV  
 QIVRDLHSWDENKLSSEFKKTSFEMTFLENQIEVSHIPNGLYYVRSIIQTDVSYPAEFLFEMTDQTV EPLVIVAK



KTDTMTTKVKLEIKVDQDHNRLLEGVGFKLVSVARDGSEKEVPLIGEYRYSSSGQVGRPLYTDKNGEIVFTNLPLGN  
YRFKEVEPLAGYAVTTLDTDVQLVDHQLVTITVVNQKLPRGNVDFMKVDGRNTSLQGAMFKVMKEESGHYTPVL  
QNGKEVVVTSKDGFRFRVEGLEYGTYLWELQAPTGYVQLTSPVSFTIGKDTRKELVTVVKNNKRPRIDVPDTGE  
ETLYILMLVAILLFGSGYYLTKKPNN

5

Orf6\_670 is a sortase. An example of an amino acid sequence of orf6\_670 is set forth in SEQ ID NO: 176.

**SEQ ID NO: 176**

MLIKMVKTKKQKRNNLLLGVVFFIGMAVMAYPLVSRLYYRVESNQIADFDKEKATLDEADIDERMKLAQAFNDS  
LNNVVSQDPWSEEMKKKGRAEYARMLEIHERMGHVEIPVIDVDLPVYAGTAEVQLQGAGHLEGTSLPIGGNSTH  
AVITAHTGLPTAKMFTDLTKLVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
HRLVLRGHRIPYVAEVEEEFIAANKLSHLYRYLFYVAVGLIVILLWIIRRLRKKKKQPEKALKALKAAARKEVKVE  
DGQQ

10

15

Orf7\_670 is a sortase. An example of an amino acid sequence of orf7\_670 is set forth in SEQ ID NO: 177.

**SEQ ID NO: 177**

VSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAFNATLKPSEILDPFTEQEKKKGVSEYANMLKVHERIG  
YVEIPAIDQEIIPMYVGTSEEILQKGAGLLEGASLPVGGENTHTVVTAHRLPTAELFSQLDKMKKGDVFYLVHLD  
QVLAYQVDQILTVEPNDFEPVLIQHGEDYATLLTCTPYMINSHRLVLRGKRIPYTAPIAERNRAVRERQFWLWL  
LLAALVMILVLSYGVYRHRIRIVKGLEKQLEEHVKG

20

Orf8\_670 is a sortase. An example of an amino acid sequence of orf8\_670 is set forth in SEQ ID NO: 178.

**SEQ ID NO: 178**

MSKAKLQKLLGYLLMLVALVIPVYCFGQMVLSLQSLGQVKGHEIFSESVTADSYQEQLQORSLDYNQRLDSQNRIVDP  
FLAEGYEVNYQVSDDPDAVYGYLSIPSLEIMEPVYLGADYHHLAMGLAHVDGTPLPVEGKGIRSVIAGHRAEPSH  
VFFRHLQDLKVGDALYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVLAFLGILFVLWKLARLLRGK

30

As discussed above, a *S. pneumoniae* AI sequence is present in the 19A Hungary 6 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences from 19A Hungary 6 are set forth below.

ORF2\_19AH is a transcriptional regulator. An example of an amino acid sequence of

ORF2\_19AH is set forth in SEQ ID NO: 187.

**SEQ ID NO: 187**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILOELQETFEELTFNLDTOQQVQLIEHHSQ  
TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISITATGYRVRQKCGLLRSVGLDLVKNQVVGPEYRIRF  
LIALLOFHFGIEIYDLNDGSMWVTHMIVQSNLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
LKNLFMYPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQLILOHTRGKHLSSKF  
KNILGNDISNSLSFLTALTFLTRTFLEGLQNLVPYNYEYHYGIESDKPLYHISKAIQVQEWMTQKIEGVIDQHR  
LYLFSLYLTETIFSSLPAIPIFIILNNQADVNLIKSIILRNFTDKVASVTGYNILISPPPSEEHLTEPLIIITTK  
EYLPYVKKQYPKGKHHFLTIALDLHVSQORLIYQTIIVDIRKEAFDKRVAMIAKKAHYLL

40

ORF3\_19AH is a cell wall surface protein. An example of an amino acid sequence of

ORF3\_19AH is set forth in SEQ ID NO: 188.

**SEQ ID NO: 188**

MKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETSIPAIGKVVIKETGEGGALLGDAVFELKNNTDGTTVSQRT  
EAQTGEAIFSNIKPGTYTLTEAQPVPVGYKPSTKQWTVEVEKNGRTTVQGEQVENREEALSQYPQTGTYPDVQTP  
YQIIKVDGSEKNGQHKALNPNPYERVIEGTLSKRIYQVNNLDDNQYGIELTVSGKTTVETKEASTPLDVVILLD  
NSNSMSNIRHNHAHRAEKAGEATRALVDKITSNPDNRVALVTYGSTIFDGSEATVEKGVADANGKILNDSALWTF  
DRTTFTAKTYNYSFLNLTSDPTDIQTIKDRIPSDAEELNKDKLMYQFGATFTQKALMTADDILTKQARPNSKKVI  
FHITDGVPTMSYPINFKYTGTTQSYRTQLNNFKAKTPNSSGILLEDFVTWSADGEHKIVRGDGESYQMFTEKPPVT

50



DQYGVHQLSLSTSMHQRALVSAGYRFTGTDLYLYWRDSILAYPFNSSTDWITNHGDPTTWYNGNMAQDGYDVF  
 TVGVGVNGDPGTDEATATRFMQSISSSPDNYTNVADPSQILQELNRYFYTIVNEKKSIENTITDPMGELIDFQL  
 GADGRFDPADYTLTANDGSSLVNNVPTGGPQNDGGLLKNKVFYDTTEKRIRVTGLYLGTGEKVTLTYNVRLNDQ  
 FVSNKFYDTNGRITLHPKEVEKNTVRDFPIPKIRDVRKYPEITIPKEKKLGEIEFIKINKNDKKPLRDAVFSLOK  
 5 QHPDYPDIYGAIDQNGTYQNVRTGEDGKLTFKNLSDGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRDVTIV  
 PQDIPAGYEFTNDKHYITNEPIPPKREYPRRTGGIGMLPFYLLIGCMMMGVLLYTRKNP

ORF4\_19AH is a cell wall surface protein. An example of an amino acid sequence of  
 ORF4\_19AH is set forth in SEQ ID NO: 189.

**SEQ ID NO: 189**

MKSINKFLTMLAALLLTASSLFSAAATVFAADNVSTAPDAVTKTLTIHKLLLSDDLKTWDTNGPKGYDGTQSSLK  
 DLTGVVAEEIPNVYFELQKYNLTGKEKENLKDDSKWTTVHGGLTTKDGLKIETSTLKGVIYRIEDRTKTTYVGP  
 NGQVLTGSKAVPALVTLPLVNNNGTVIDAHVFPKNSYNKPVVDKRIADTLNNDQNGLSIGTKIPYVNTTIPSN  
 ATFATSFWSDEMTEGLTYNEDVTITLNNVAMDQADYEVTGKXNGFNKLTEAGLAKINGKDADQKIQITYSATLN  
 15 SLAVADIPESNDITYHYGNHGDHGNTPKPTKPNNGQITVTKTWDSQPAPEGVKATVQLVNAKTGEKVGAPVELSE  
 NNWYTWSGLDNSIEYKVEEEYNGYSAEYTVESKGLGVKNWKNPAPINPEEPRVKTYGKKFVKVDQKDTRLE  
 NAQFVVKKADSNKYIAFKSTAQQAADKAAATAKQKLDAVAAYTNAADKQAAQALVDQAQQEYNVAYKEAKFGY  
 VEVAGKDEAMVLTSNTDGQFQISGLAAGTYKLEEIKAPEGFAKIDDFEVVGAGSWNQGEFNYLKDVQKNDATKV  
 VNKKITIPQTGGIGTIIFAVAGAAIMGIAVYAYVKNNKDEDQLA

ORF5\_19AH is a cell wall surface protein. An example of an amino acid sequence of  
 ORF5\_19AH is set forth in SEQ ID NO: 190.

**SEQ ID NO: 190**

MTMQKMQKMISRIFFVMALCFSLVWGAHAVQAQEDHTLVLQLENYQEVVSQLP SRDGHRLQVWKLDDSYSDRV  
 25 QIVRDLHSWDENKLSSFKKTSFEMTFLENQIEVSHIPNGLYYVRSIIQTDVASYPAEFLFEMTDQTVPLVIVAK  
 KTDMTTKVKLIKVDQDHNRLLEGVGFKLVSVDGSEKEVPLIGEYRYSSSGQVGRITLYTDKNGEIVTNLPLGN  
 YRFKEVEPLAGYAVTTLDTDVQLVDHQLVTITVNVNQLPRGNVDFMKVDGRNTNTSLQGAMFKVMKEESGHYTPVL  
 QNGKEVVVTSGKDGFRFRVEGLEYGTYLWELQAPTGYVQLTSPVSFTIGKDTRKELTVVKNKRPRIDVPDTGE  
 ETLYILMLVAILLFGSGYYLTCKPNN

ORF6\_19AH is a putative sortase. An example of an amino acid sequence of ORF6\_19AH is  
 set forth in SEQ ID NO: 191.

**SEQ ID NO: 191**

MLIKMKVTKKQKRNLLLGVVFFIGMAVMAYPLVSRLYYRVE SNQQIADFDKEKATLDEADI DERMKLAQAFNDS  
 35 LNNVSGDPWSEEMKKKGRAEYARMLEIHERMGHVEIPVIDVDLPVYAGTAEVVLQOGAGHLEGTSLPIGGNSTH  
 AVITAHTGLPTAKMFTDLTKLVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
 HRLVLRGHRIPYVAEVEEEFIAANKLSHLYRYLFYVAVGLIVILLWIIRRLRKKKKQPEKALKALKAAARKEVKVE  
 DGQQ

ORF7\_19AH is a putative sortase. An example of an amino acid sequence of ORF7\_19AH is  
 set forth in SEQ ID NO: 192.

**SEQ ID NO: 192**

MDNSRRSRKKGTKKKKHPLILLILFLVGFAVAIYPLVSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAF  
 NATLKPSEILDPFTEQEKKGVSSEYANMLKVHERIGYVEIPAIQDQEIIPMYVGTSEEILQKGAGLLEGASLPVGGE  
 45 NTHTVVTAHRLPTAELFSQLDKMKKGDVFYLVLDQVLAYQVDQILTVEPNDFEPVLIQHGEDYATLLTCTPYM  
 INSHRLVLRGKRIPYTAPIAERNRAVRERGQFWLWLLAALVMILVLSYGVYRHRIRIVKGLEKQLEEHVKG

ORF8\_19AH is a putative sortase. An example of an amino acid sequence of ORF8\_19AH is  
 set forth in SEQ ID NO: 193.

**SEQ ID NO: 193**

MSKAKLQKLLGYLLMLVALVIPVYCFGQMVLSLQGVKGHEIFSESVTADSYQEQLQRLSDYNQRLDSQNRIVDP  
 FLAEGYEVNYQVSDDPDVYGYLSIPSLEIMEPVYLGADYHHLAMGLAHVDGTPLPVEGKGIRSVIAGHRAEPSH  
 VFFRHLQKLVGDALYYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
 55 YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAFMGILFVLWKLARLLRGK



As discussed above, a *S. pneumoniae* AI sequence is present in the 6B Finland 12 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences from 6B Finland 12 are set forth below.

ORF2\_6BF is a transcriptional regulator. An example of an amino acid sequence of

5 ORF2\_6BF is set forth in SEQ ID NO: 194.

**SEQ ID NO: 194**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDLTQQVQLIEHHS HQ  
TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISIAATGYRVRQKCGLLRSVGLDLVKNQVVGPEYRIRF  
LIALLQFHFGIEIYDLNDGSMWVTHMIVQNSQLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
10 LKNLFMYPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQLILQHTRGKHLSSKF  
KNILGNDISNLSFLTALTFLTRTFGLQNLVPYNYEYHYGIESDKPLYHISKAIQEWMTQKIEGVIDQHR  
LYLFSLYLTETIFSSLPAIPFIILNNQADVNLIKSIIILRNFTDKVASVTGYNILISPPPSEHLTEPLIIITTK  
EYLPYVKKQYPKGKHHFLTIALDLHVSQORLIYQTIIVDIRKEAFDKRVAMIAKKAHYLL

15 ORF3\_6BF is a cell wall surface protein. An example of an amino acid sequence of

ORF3\_6BF is set forth in SEQ ID NO: 195.

**SEQ ID NO: 195**

MKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETSIPAIGKVVIKETGEGGALLGDAVFELKNNTDGTTVSQRT  
EAQTGEAIFSNIKPGTYTLTEAQQPVGYKPKSTKQWTVVEVEKNGRTTVQGEQVENREEALSDQYPQTGTYPDVQTP  
20 YQIIKVDGSEKNGQHKALNPYPYERVIPEGTLISKRIYQVNNLDDNQYGIELTVSGKTTVETKEASTPLDVVILLD  
NSNSMSNIRHNHAHRAEKAGEATRALVDKITSNPDNRVALVTYGSTIFDGSEATVEKGVADANGKILNDSALWTF  
DRTTFTAKTYNYSFLNLTSDPTDIQTIKDRIPSDAEELNKDKLMYQFGATFTQKALMTADDILTQARPNSKKVI  
FHITDGVPTMSYPINFKYTGTTQSYRTQLNNEFKAKTPNSSGILLEDFVTSADGEHKIVRGDGEYQMFTEKPKVT  
DQYGVHQILSITSMEQRAKLVSAGYRFYGTDLYLWRDSILAYPFNSSTDWITNHGDPTTWYYNGNMAQDGYDVF  
25 TVGVGVNGDPGTDEATATRFMQSISSSPDNYTNVADPSQILQELNRYFYTIVNEKKSIEGTITDPMGELIDFQL  
GADGRFDPADYTLTANDGSSLVNNVPTGGPQNDGGLLKNKAVFYDTTEKRIRVTGLYLGTGEKVTLTYNVRLNDQ  
FVSNKFYDTNGRTTLHPKEVEKNTVRDFPIPKIRDVRKYPEITIPKEKKLGEIEFIKINKNDKKPLRDAVFSLQK  
QHPDYPDIYGAIDQNGTYQNVRTGEDGKLTFKNLSDGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRDVTISIV  
PQDIPAGYEFTNDKHYITNEPIPPKREYPRRTGGIGMLPFYILGCMGMMGGVLLYTRKHP

30 ORF4\_6BF is a cell wall surface protein. An example of an amino acid sequence of

ORF4\_6BF is set forth in SEQ ID NO: 196.

**SEQ ID NO: 196**

MKSINKFLTMLAALLLTASSLFSAAVFAADNVSTAPDAVTKTLTIHKLLLSEDDLKTWDTNGPKGYDGTQSSSLK  
35 DLTGVVAEEIPNVYFELQKYNLTGKEKENLKDSSKWTTVHGGTLTKDGLKIETSTLKGVIYRIREDRTKTTYVGP  
NGQVLTGSKAVPALVTLPLVNNNGTVIDAHVFPKNSYNKPVVDKRIADTLNNDQNGLSIGTKIPYVVNTTIPSN  
ATFATSFWSDTEGLTYNEDVTITLNNVAMDQADYEVTKGNNGFNKLTEAGLAKINGKDADQKIQITYSATLN  
SLAVADIPESENDITYHYGNHQDHGNTPKPTKPNNGQITVTKTWDSQPAPEGVKATVQLVNAKTGEKVGAPVELSE  
NNWYTWGLDNSIEYKVEEEYNGYSAEYTVESKGLGVKNWKNPAPINPEEPRVKTYGKKFVKVDQKDRLE  
40 NAQFVVKKADS NKYIAFKSTAQQAADEKAAATAKQKLDAVAAYTNAADKQAAQALVDQAQQEYNVAYKEAKFGY  
VEVAGKDEAMVLTSTNDGQFQISGLAAGTYKLEEIKAPEGFAKIDDFEVVVGAGSWNQGEFNYLKDVQKNDATKV  
VNKKITIPQTGGIGITIIFAVAGAAIMGIAVYAYVKNKDEDQLA

ORF5\_6BF is a cell wall surface protein. An example of an amino acid sequence of

45 ORF5\_6BF is set forth in SEQ ID NO: 197.

**SEQ ID NO: 197**

MTMQKMQMISRIFFVMALCFSLVWGAHAVQAQEDHTLVQLQENYQEVVSQVPSRDGHRQLQVWKLDDSYSDDRV  
QIVRDLHSWDENKLSFFKTSFEMTFLENQIEVSHIPNGLYYVRSIIQTDAVSYPAEFLFEMTDQTVPLVIVAK  
KTDMTTKVKLIKVDQDHNRLGEGVGFKLVSARDGSEKEVPLIGEYRYSSSGQVGRITLYTDKNGEIVFTNLPLGN  
50 YRFKEVEPLAGYAVTTLDTDVQLVDHQLVTITVVNQKLPRGNVDFMKVDGRTNTSLQGAMFKVMKEESGHYTPVL  
QNGKEVVVTSKDGGRFRVEGLEYGTYLWELQAPTGYVQLTSPVSFTIGKDRKELVTVVKNKRPRIIDVPDTGE  
ETLYILMLVAILLFGSGYYLTCKPNN



ORF6\_6BF is a putative sortase. An example of an amino acid sequence of ORF6\_6BF is set forth in SEQ ID NO: 198.

**SEQ ID NO: 198**

5 MLIKMKVTKKQKRNNLLLGVVFFIGMAVMAYPLVSRLYYRVESNQIADFDKEKATLDEADI DERMKLAQAFNDS  
LNNVSGDPWSEEMKKKGRAEYARMLEIHERMGHVEIPVIDVDLPVYAGTAEV LQQGAGHLEGTSLPIGGNSTH  
AVITAHTGLPTAKMFTDLTKLKVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
HRLLVRGHRI PYVAEVEEEEFIAANKLSHLYRYLFYVAVGLIVILLWIIRRLRKKKKQPEKALKALKAARKEVKVE  
DGQQ

10 ORF7\_6BF is a putative sortase. An example of an amino acid sequence of ORF7\_6BF is set forth in SEQ ID NO: 199.

**SEQ ID NO: 199**

15 MDNSRRSRKKGTKKKKHPLILLIFLVGFAVAIYPLVSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAF  
NATLKPSEILDPFTEQEKKKGVSEYANMLKVHERIGYVEI PAIDQEIPMYVGTSEEILQKGAGLLEGASLPVGGE  
NTHTVTAHRGLPTAELFSQLDKMKKGDVFYLVLDQVLAYQVDQILTVEPNDFEPVLIQHGEDYATLLTCTPYM  
INSHRLLVRGKRIPYTAPIAERNRAVRERGGQFWLWLLLAALVMILVLSYGVYRHRIRIVKGLEKQLEEHVKG

ORF8\_6BF is a putative sortase. An example of an amino acid sequence of ORF8\_6BF is set forth in SEQ ID NO: 200.

**SEQ ID NO: 200**

20 MSKAKLQKLLGYLLMLVALVIPVYCFGQMVLQSLGQVKGHEIFSESVTADSYQEQLQSLDYNQRLDSQNRIVDP  
FLAEGYEVNYQVSDDPDAVYGYSIPSLEIMEPVYLGADYHHLAMGLAHVDGTPLPVEGKGIRSVIAGHRAEPSH  
VFFRHLDQLKVGDALYYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAFGLFVLWKLARLLRGK

25 As discussed above, a *S. pneumoniae* AI sequence is present in the 6B Spain 2 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences from 6B Spain 2 are set forth below.

ORF2\_6BSP is a transcriptional regulator. An example of an amino acid sequence of ORF2\_6BSP is set forth in SEQ ID NO: 201.

**SEQ ID NO: 201**

30 MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDTQQVQLIEHHSHQ  
TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISIAATGYRVRQKCGLLRSVGLDLVKNQVVGPEYRIRF  
LIALLQHFHFGIEIYDLNDGSMDWVTHMIVQSNLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
35 LKNLFMYPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKTHTIQLILQHTRGKHLLSKF  
KNILGNDISNLSFLTALTFLTRTFGLQNLVPYNYEYHYGIESDKPLYHISKAIVQEWMTQKIEGVIDQHR  
LYLFSLYLTETIFSSLPAIPFIILNNQADVNLIKSIILRNFTDKVASVTGYNILISPPPSEEHLTEPLIIITTK  
EYLPYVKKQYPKGGKHFFLTIALDLHVSQQRLIYQTIVDIRKEAFDKRVAMIAKKAHYLL

ORF3\_6BSP is a cell wall surface protein. An example of an amino acid sequence of

40 ORF3\_6BSP is set forth in SEQ ID NO: 202.

**SEQ ID NO: 202**

MKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETS PAIGKVVIKETGEGGALLGDAVFELKNNTDGTTVSQR  
EAQTGEAIFSNIKPGTYTLTEAQPVGYPSTKQWTVVEVEKNGRRTTVQGEQVENREEALSDQYPQTGTYPDVQTP  
YQIIKVDGSEKNGQHKALNPYPYERVIPEGTL SKRIYQVNNLDDNQYGIELTVSGKTTVETKEASTPLDVILLD  
45 NSNSMSNIRHNHAHRAEKAGEATRALVDKITSNPDNRVALVTYGSTIFDGSEATVEKGVADANGKILNDSALWTF  
DRTTFTAKTYNYSFLNLTSDPTDIQTIKDRIPSDAEELNKDKL MYQFGATFTQKALMTADDILTKQARPNSKKVI  
FHITDGVPTMSYPINFKYTGTTQSYRTQLN NFKA KTPNSSGILLEDFVTWSADGEHKIVRGDGE SYQMFTKKPVT  
DQYGVHQILSITSMEQRAKLVSAGYRFYGTDL YLYWRDSILAYPFNSSTDWITNHGDPTTWYNGNMAQDGYDVF  
TVGVGVNGDPGTDEATATRFMQSISSSPDNYTNVADPSQILQELNRYFYTIVNEKKS IENG TITDPMGELIDFQL  
50 GADGRFDPADYTLTANDGSSLVNNVPTGGPQNDGGLLKN AKVFYDTTEKRIRVTGLYLGTGEKVTLTYNVRLNDQ  
FVSNKFYDTNGRTTLHPKEVEKNTVRDFPIPKIRDVRKYPEITIPKEKKLGEIEFIKINKNDKKPLRDAVFS LQK  
QHPDYPDIYGAIDQNGTYQNVRTGEDGKLTFKNLSDGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRDVT SIV  
PQDIPAGYEFTNDKHYITNEPIPPKREYPR TGGIGMLPFYLGMMMGVLLYTRKHP



ORF4\_6BSP is a cell wall surface protein. An example of an amino acid sequence of

ORF4\_6BSP is set forth in SEQ ID NO: 203.

**SEQ ID NO: 203**

5 MKSINKFLTMLAALLLTASSLFSAAATVFAADNVSTAPDAVTKTLTIHKLLLSDDLKTWDTNGPKGYDGTQSSLK  
DLTGVAEEIPNVYFELQKYNLTGKEKENLKDDSKWTTVHGGLTTKDGLKIETSTLKGVIYRIREDRTKTTYVGP  
NGQVLTGSKAVPALVTLPLVNNNGTVIDAHVFPKNSYNKPVDKRIADTLNNDQNGLSIGTKIPYVNTTIPSN  
ATFATSFWSDEMTEGLTYNEDVTITLNNVAMDQADYEVTGKNGGFLNKLTEAGLAKINGKDADQKIQTYSATLN  
SLAVADIPESNDITYHYGNHQDHGNTPKPTKPNNGQITVTKTWDSQPAPEGVKATVQLVNAKTGEKVGAPVELSE  
10 NNWYTWGSLDNSIEYKVEEEYNGYSAEYTVESKGLGVKNWKNPAPINPEEPRVKTYGKKFVKVDQKDTRLE  
NAQFVVKKADSNKYIAFKSTAQQAADEKAAATAKQKLDAVAAYTNAADKQAAQALVDQAQQEYNVAYKEAKFGY  
VEVAGKDEAMVLTSNTDGQFQISGLAAGTYKLEEIKAPEGEFAKIDDFEVVGAGSWNQGEFNYLKDVQKNDATKV  
VNKKITIPQTGGIGTIIFAVAGAAIMGIAVYAYVKNKDEDDQLA

ORF5\_6BSP is a cell wall surface protein. An example of an amino acid sequence of

15 ORF5\_6BSP is set forth in SEQ ID NO: 204.

**SEQ ID NO: 204**

MTMQKMQMISRIFFVMALCFSLVWGAHAVQAQEDHTLVLQLENYQEVVSQLP SRDGHRLQVWKLDDSYSDRRV  
QIVRDLHSWDENKLSSFKKTSFEMTFLENQIEVSHIPNGLYYVRSIIQTDVSYPAEFLFEMTDQTEPLVIVAK  
KTDTMTTKVKLIKVDQDHNRLGEGVGFKLVSVDGSEKEVPLIGEYRYS SSGQVGR TLYTDKNGEIVTNLPLGN  
20 YRFKEVEPLAGYAVTTLDTDVQLVDHQLVTITVNVNQLPRGNVDFMKVDGRTNTSLQGAMFKVMKEESGHYTPVL  
QNGKEVVVTSGKDGRFRVEGLEYGTYLWELQAPTGYVQLTSPVSFTIGKDTRKELVTVVKNKRPRIDVPDTGE  
ETLYILMLVAILLFGSGYYLTKKPNN

ORF6\_6BSP is a putative sortase. An example of an amino acid sequence of ORF6\_6BSP is

25 set forth in SEQ ID NO: 205.

**SEQ ID NO: 205**

MLIKMVKTKKQKRNNLLLGVVFFIGMAVMAYPLVSRLYYRVESNQQIADF DKEKATLDEADI DERMKLAQAFNDS  
LNNVVS GDPWSEEMKKKGRAEYARMLEIHERMGHVEIPVIDVDLPVYAGTAEV LQQGAGHLEGTSLPIGGNSTH  
AVITAHTGLPTAKMFTDLTKLVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
30 HRLLVRGHRI PYVAEVEEEFIAANKLSHLRYLRYFVAVGLIVILLWII RRLRKKKKQPEKALKALKAAARKEVKVE  
DGQQ

ORF7\_6BSP is a putative sortase. An example of an amino acid sequence of ORF7\_6BSP is  
set forth in SEQ ID NO: 206.

35 **SEQ ID NO: 206**

MDNSRRSRKKGTKKKKHPLILLIFLVGFVAIYPLVSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAF  
NATLKPSEILDPFTEQEKKKGVS EYANMLKVHERIGYVEIPAI DQEI PMYVGTSEEILQKGAGLLEGASLPVGGE  
NTHTVVTAHRGLPTAELFSQLDKMKKGDFYLVHLDQVLAYQVDQILTVEPNDFEPVLIQHGEDYATLLTCTPYM  
40 INSHRLLVRGKRIPYTAPIAERNRAVRERGQFWLWLLAALVMILVLSYGVYRHRRI VKGLEKQLEEHVKG

ORF8\_6BSP is a putative sortase. An example of an amino acid sequence of ORF8\_6BSP is  
set forth in SEQ ID NO: 207.

**SEQ ID NO: 207**

MSKAKLQKLLGYLLMLVALVIPVYCFGQMVLSLQSLGQVKGHEIFSESVTADSYQEQLQRLDYNQRLDSQNRIVDP  
45 FLAEGYEVNYQVSDDPDVYGYLSIPSLEIMEPVYLGADYHHLAMGLAHVDGTPLPVEGKGIRSVIAGHRAEPSH  
VFFRHLQKLVGDALYYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAF LGILFVLWKLARLLRGK

As discussed above, a *S. pneumoniae* AI sequence is present in the 9V Spain 3 *S. pneumoniae*  
50 genome. Examples of *S. pneumoniae* AI sequences from 9V Spain 3 are set forth below.

ORF2\_9VSP is a transcriptional regulator. An example of an amino acid sequence of  
ORF2\_9VSP is set forth in SEQ ID NO: 208.



**SEQ ID NO: 208**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDTQQVQLIEHHSHQ  
 TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISISATGYRVRQKCGLLRSVGLDLVKNQVVGPEYRIRF  
 LIALLOFHFGIEIYDLNDGSMWVTHMIVQSNLSHLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
 LKNLFMYPILEHMCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQLILQHTRGKHLSSKF  
 KNILGNDISNLSLFTALTFLTRTFLFGLQNLVPYNNYEHYGIESDKPLYHISKAIVQEWMTQKIEGVIDQHR  
 LYLFSLYLTETIFSSLPAPIFIILNNQADVNLIKSIILRNFTDKVASVTGYNILISPPPSEEHLTEPLIIITTK  
 EYLPYVKKQYPKGKHHFLTIALDLHVSQORLIYQTIIVDIRKEAFDKRVAMIAKKAHYLL

ORF3\_9VSP is a cell wall surface protein. An example of an amino acid sequence of  
 ORF3\_9VSP is set forth in SEQ ID NO: 209.

**SEQ ID NO: 209**

MKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETSIPAIGKVVIKETGEGGALLGDAVFELKNNTNGTTVSQRT  
 EAQTGEAIFSNIKPGTYTLTEAQPVGYPKSTKQRTVEVEKNGRTTVQGEQVENREEALSDQYPQTGTYPDVQTP  
 YQIIKVDGSEKNGQHKALNPYPYERVIPEGTLISKRIYQVNNLDDNQYGIELTVSGKTVYERKDKSVPLDVVILL  
 NSNSMSNIRNKNARRAERAGEATRLIDKITSDPENRVALVTYASTIFDGTEFTVEKGVADKNGKRLNDSLFWNY  
 DQTSFTTNTKDYSLKLTNDKNDIVELKNKVPTAEADHDGNRLMYQFGATFTQKALMKADEILTQQARQNSQKVI  
 FHITDGVPTMSYPINFHATFAPSYQNQLNAFFSKSPNKDGILLSDFITQATSGEHTIVRGDGSYQMFDTKTVY  
 EKGAPAAFPVKPEKYSEMKAAGYAVIGDPINGGYIWLNWRESILAYPFNSNTAKITNHGDPTRWYNGNIAPDGY  
 DVFTVGIGINGDPTDEATATSEMQSISSKPENYTNVTDTTKILEQLNRYFHTIVTEKKSIENGITDPMGELID  
 LQLGTDGRFDPADYTLTANDGSRLENGQAVGGPQNDGGLLKNAKFYDTEKRIKRVGLYLGTGEKVTLTYNVRL  
 NDQFVSNKFYDTNGRTTLHPKEVEKNTVRDFPIPKIRDVRKYPATITAKEKKLGEIEFIKINKNDKKPLRDVFS  
 LQKQHPDYPDIYGAIQNGTYQNVRTGEDGKLTFFKNLSDGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRDVT  
 SIVPQDIPAGYEFTNDKHYITNEPIPPKREYPRGTGGIGMLLFYLGCMGGVLLYTRKHP

ORF4\_9VSP is a cell wall surface protein. An example of an amino acid sequence of  
 ORF4\_9VSP is set forth in SEQ ID NO: 210.

**SEQ ID NO: 210**

MKSINKFLTMLAALLLTASSLFSAAATVFAAGTTTTSVTVHKLATDGDMDKIANELETGNYAGNKVGLPANAKE  
 IAGVMFVWTNTNNEIIDENGQTLGVNIDPQTFKLSGAMPATAMKKLTEAEGAKFNTANLPAAKYKIYEIHSLSY  
 VGEDGATLTGSKAVPIEIELPLNDVDAHVYPKNTEAKPKIDKDFKGANPDTPRVKDTFVNHQVGDVVEYEIV  
 TKIPALANYATANWSDRMTEGLAFNKGTVKVTVDVALEAGDYALTEVATGFDLKLTDAGLAKVNDQNAEKTVKI  
 TYSATLNDKAIVEVPESNDVTFNYGNNPDHGNTPKPNKPNENGDLTLTKTWVDATGAPIPAGAEATFDLVNAQTG  
 KVVQTVTLTTDKNTVTVNGLDKNTYKFFVERSISGYSADYQEITTAGETIAVKNWKDENPKPLDPTPKVVTYGGK  
 FVKVNDKDNRLAGAEFVIANADNAGQYLARKADKVSQEEKQLVVTTKDALDRAVAAYNALTAQQQTQOEKEKVDK  
 AQAAYNAAVIAANNAFEWVADKDNENVVKLVSDAQGRFEITGLLAGTYYLEETKQPAGYALLTSRQKFEVTATSY  
 SATGQGIETAGSGKDDATKVVNKKITIPQTGGIGITIIFAVAGAVIMGIAVYAVKNNKDEDQLA

ORF5\_9VSP is a cell wall surface protein. An example of an amino acid sequence of  
 ORF5\_9VSP is set forth in SEQ ID NO: 211.

**SEQ ID NO: 211**

MTMQKMQKMQKMQKMQKMISRIFFVMALCFSLVWGAHAVQAQEDHTLVQLQENYQEVVSQLP SRDGHRLQVW  
 KLDDSYSYDNRVQIVRDLHSWDENKLSSEFKKTSFEMTFLENQIEVSHIPNGLYYVRSIIQTDAVSYPAEFLFEMT  
 DQTVETPLVIVAKKADTVTTKVKLKVDQDHNRLLEGVGFKLVS VARDGSEKEVPLIGEYRYSSSGQVGR TLYTDKN  
 GEIVVTNLPLGTYRFKEVEPLAGYTVTTMDTDVQLVDHQLVTITVVNQKLPRGNVDFMKVDGRTNTSLQGAMFKV  
 MKEENGHYTPVLQNGKEVVVASGKDGRFRVEGLEGYTYLWELQAPTGYVQLTSPVSFTIGKDRKELVTVVKNN  
 KRPRIDVPDTGEETLYILMLVAILLFGSGYYLTCKTNN

ORF6\_9VSP is a putative sortase. An example of an amino acid sequence of ORF6\_9VSP is  
 set forth in SEQ ID NO: 212.

**SEQ ID NO: 212**

MLIKMAKTKKQKRNNLLLGVVFFIGIAVMAYPLVSRLYYRVESNQQIADFDKEKATLDEADI DERMKLAQAFNDS  
 LNNVSGDPWSEEMKKKGRAEYARMLEIHERMGHVEIPAIDVDLPVYAGTAEV LQQGAGHLEG TSLPIGGNSTH  
 AVITAHTGLPTAKMFTDLTKLVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
 HRLLRGHRIPYVAEVEEEFIAANKLSHLYRYLFYVAVGLIVILLWIIIRLRKKKRQSERALKALKEATKEVKVE  
 DE



PCT/US05/27239

ORF7\_9VSP is a putative sortase. An example of an amino acid sequence of ORF7\_9VSP is set forth in SEQ ID NO: 213.

**SEQ ID NO: 213**

5 MSKSRYSRKKS VKKKKNPFILLIFLVGLAVAMYPLVSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAF  
NATLKPSEILDPFTEQEKKKGVSEYANMLKVHERIGYVEIPAIQEI PMYVGTSEEILQKGAGLLEGASLPVGGE  
NTHTVVTAHRGLPTAELFSQLDKMKKGDIFYLHVLDQVLAYQVDQIVTVEPNDFEPVLIQHGEDYATLLTCTPYM  
INSHRLLVRGKRIPYTAPIAERNRAVRERGGQFWLWLLLGAMAVILLLLYRVYRNRRIVKGLEKQLEGRHVKD

10 ORF8\_9VSP is a putative sortase. An example of an amino acid sequence of ORF8\_9VSP is set forth in SEQ ID NO: 214.

**SEQ ID NO: 214**

MSRTKL RALLGYLLMLVACLIPIYCFGQMVLQSLGQVKGHATFVKSM TTEMYQEQQNHSLAYNQRLASQNRI VDP  
FLAEGYEVNYQVSDDPDAVYGYLSIPSLEIMEPVYLGADYHHLGMGLAHVDGTPLPLDGTGIRSVIAGHRAEPSH  
15 VFRHLDQLKVG DALYYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAF LGILFVLWKLARLLRGK

As discussed above, a *S. pneumoniae* AI sequence is present in the 14 CSR 10 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences from 14 CSR 10 are set forth below.

20 ORF2\_14CSR is a transcriptional regulator. An example of an amino acid sequence of ORF2\_14CSR is set forth in SEQ ID NO: 215.

**SEQ ID NO: 215**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLSQSKLLSILQELQETFEELTFNLDTQQVQLIEHHS HQ  
TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYIS IATGYRVRQKCGLLRSVGLDLVKNQVVGPEYRIRF  
25 LIALLOFHFGIEIYDLNDGSM DWVTHMIVQSNSQLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
LKNLFMYPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQILILQHTRGKHL LSKF  
KNILGNDISNSLSFLTALTFLTRTFLEGLQNLVPYYNYEHYGI ESDKPLYHISKAIVQEWMT EQKIEGVIDQHR  
LYLFSLYLTETIFSSLPAPIFIILNNQADVNLKSIILRNFTDKVASVTGYNILISPPPSEEHLTEPLIIITTK  
EYLPYVKKQYPKGKHHFLTIALDLHVSQQRLIYQTIVDIRKEAFDKRVAMI AKKAHYLL

30 ORF3\_14CSR is a cell wall surface protein. An example of an amino acid sequence of ORF3\_14CSR is set forth in SEQ ID NO: 216.

**SEQ ID NO: 216**

MKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETS PAIGKVVIKETGEGGALLGDAVFELKNNTDGT TVSQRT  
35 EAQTGEAIFSNIKPGTYTLTEAQPPVGYK PSTKQWTVEVEKNGRTTVQGEQVENREEALSDQYPQTGTPDVQTP  
YQIIKVDGSEKNGQH KALNPNPYERVIPEGT LSKRIYQVNNLDDNQYGIELTVSGKTTVETKEASTPLD VVILLD  
NSNSMSNIRHNHAHRAEKAGEATRALVDKITSNPDNRVALVTY GSTIFDGSEATVEKGVADANGKILNDSALWTF  
DRTTFTAKTYNYSFLNLTSDPTDIQTIKDRIPSDAEELNKDKLMYQFGATFTQKALMTADDILTKQARP NSKKVI  
FHITDGVPTMSYPINFKYTGTTQSYRTQLNNFKAKTPNSSGILLEDFVTWSADGEHKIVRGDGESYQMFTK KPV  
40 DQYGVHQILSITSMEQRAKLVSAGYRFYGTDL YLYWRDSILAYPNSSTDWITNHGDPTTWYNGNMAQDGYDVF  
TVGVGVNGDPGTDEATATRFMQSISSSPDNYTNVADPSQILQELNRYFYTIVNEKKSIENG TITDPMGELIDFQL  
GADGRFDPADYTLTANDGSSLVNNVPTGGPQNDGGLLKN AKVFYDTTEKRIRVTGLYLGTGEKVTLTYNVR LNDQ  
FVSNKFYDTNGR TTHLPKEVEKNTVRDFPIPKIRDVRKYPEITIPKEKKLGEIEFIKINKNDKKPLRDAV FSLQK  
QHPDYPDIYGAIDQNGTYQNVRTGEDGKLTFKNLS DGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRD VTSIV  
45 PQDIPAGYEFTNDKHYITNEPIPPKREYPR TGGIGMLPFYLLIGCMMMGGVLLYTRKHP

ORF4\_14CSR is a cell wall surface protein. An example of an amino acid sequence of ORF4\_14CSR is set forth in SEQ ID NO: 217.

**SEQ ID NO: 217**

50 MKSINKFLTMLAALLLTASSLFS AATVFAADNVSTAPDAVTKTLTIHKLLS EDDLKTWDTNGPKGYDGTQSS LK  
DLTGVAEEIPNVYFELQKYNLTDGKEKENLKDDSKWTTVHGGLTTKDGLKIETSTLKG VYRIREDRTKTTYVGP  
NGQVLTGSKAVPALVTLPLVNNNGTVIDAHVFPKNSYNKPVVDKRIADTLNNDQNGLSIGTKIPYV VNTTIPSN  
ATFATSFWSDEMTEGLTYNEDVTITLNNVAMDQADYEVT KGNNGFNKLKTEAGLAKINGKDADQKI QITYSATLN  
SLAVADIPESNDITYHYGNHQDHGNTPKPTKPNNGQITVTKTWDSQPAPEGVKATVQLVNAKTGEKVGAPV ELSE



~~NNWLYTWSGLDNSTELKVEEENGYBAETVESK~~KGKLGVKWNKDNNPAPINPEEPRVKTYGKKFVKVDQKDTRLE  
 NAQFVVKKADSNKYIAFKSTAQQAADKAAATAKQKLDAAVAAYTNAADKQAAQALVDQAQQEYNVAYKEAKFGY  
 VEVAGKDEAMVLTSTNDGQFQISGLAAGTYKLEEIKAPEGFAKIDDFEVVVGAGSWNQGEFNYLKDVQKNDATKV  
 VNKKITIPQTGGIGTIIFAVAGAAIMGLAVYAYVKNKDEDQLA

ORF5\_14CSR is a cell wall surface protein. An example of an amino acid sequence of ORF5\_14CSR is set forth in SEQ ID NO: 218.

**SEQ ID NO: 218**

MTMQKMQMISRIFFVMALCFSLVWGAHAVQAQEDHTLVQLQENYQEVVSQLP SRDGHRLQVWKLDDSYSDDRV  
 QIVRDLHSWDENKLSSEFKKTSFEMTFLENQIEVSHIPNGLYYVRSIIQTDVSYPAEFLFEMTDQTV EPLVIVAK  
 KTDMTTKVKLIKVDQDHNRLGEGVGFKLVS VARDGSEKEVPLIGEYRYSSSGQVGR TLYTDKNGEIFVTNLPLGN  
 YRFKEVEPLAGYAVTTLDTDVQLVDHQLVTITVVNQKLPRGNVDFMKVDGR TNTSLQGAMFKVMKEESGHYTPVL  
 QNGKEVVVTSGKDGFRFVEGLEYGTYLWELQAPTGYVQLTSPVSFTIGKDTRKELTVVVKNNKRPRIDVPDTGE  
 ETLYILMLVAILLFGSGYYLTKKPNN

ORF6\_14CSR is a putative sortase. An example of an amino acid sequence of ORF6\_14CSR is set forth in SEQ ID NO: 219.

**SEQ ID NO: 219**

MLIKMKVTKKQKRNNLLLG VVFFIGMAVMAYPLVSRLYYRVESNQIADFDKEKATLDEADI DERMKLAQAFNDS  
 LNNVVS G DPWSEEMKKKGRAEYARMLEIHERM GHVEIPVIDVDLPVYAGTAEV LQQGAGHLEGTSLPIGGNSTH  
 AVITAHTGLPTAKMFTDLTKLVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
 HRLLRGHRIPYVAEVEEEFIAANKLSHLYRYLFYVAVGLIVILLWIIRRLRKKKKQPEKALKALKAAARKEVKVE  
 DGQQ

ORF7\_14CSR is a putative sortase. An example of an amino acid sequence of ORF7\_14CSR is set forth in SEQ ID NO: 220.

**SEQ ID NO: 220**

MDNSRRSRKKGTKKKKHPLILLIIFLVGF AVAIYPLVSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAF  
 NATLKPSEILD PFTEQEKKKG VSEYANMLKVHERIGYVEIPAI DQEIPMYVGTSEEILQKGAGLLEGASLPVGGE  
 NTHTVVTAHRGLPTAELFSQLDKMKKG DVFYLVLDQVLAYQVDQILTVEPNDFEPVLIQHGEDYATLLTCTPYM  
 INSHRLLVRGKRIPYTAPIAERNRAVRERGQFWLWLLLAALVMILVLSYGVYRHRIRIVKGLEKQLEEHVKG

ORF8\_14CSR is a putative sortase. An example of an amino acid sequence of ORF8\_14CSR is set forth in SEQ ID NO: 221.

**SEQ ID NO: 221**

MSKAKLQKLLGYLLMLVALVIPVYCFGQMV LQSLGQVKGHEIFSESVTADSYQEQLQ RSLDYNQRLDSQNRIVDP  
 FLAEGYEVNYQVSDDPDAVYGYLSIPSLEIMEPVYLGADYHHLAMGLAHVDGTPLPVEGKGIRSVIAGHRAEPSH  
 VFFRHLQDKVGDALYYDNGQEIVEYQMM DTEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
 YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVLAFLGILFVLWKLARLLRGK

As discussed above, a *S. pneumoniae* AI sequence is present in the 19F Taiwan 14 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences from 19F Taiwan 14 are set forth below.

ORF2\_19FTW is a transcriptional regulator. An example of an amino acid sequence of ORF2\_19FTW is set forth in SEQ ID NO: 222.

**SEQ ID NO: 222**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDTQQVQLIEHHS HQ  
 TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYIS IATGYRVRQKCGLLRSVGLDLVKNQVVGPEYRIRF  
 LIALLQFHFGIEIYDLNDGSM DWVTHMIVQSNSQLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
 LKNLFMPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQLILQHTRGKHL LSKF  
 KNILGNDISNSLSFLTALTFLTRTFLFGLQNLVPYNYEYHYGIESDKPLYHISKAIVQEWMT EQKIEGVIDQHR  
 LYLFSLYLTETIFSSLPAIPIFIILNNQADVNLIKSIILRNFTDKVASVTGYNILISPPPSEEHLTEPLIIITTK  
 EYLPYVKKQYPKGKHHFLTIALDLHVSQQRLIYQTIVDIRKEAFDKRVAMIAKKAHYLL



PC/T/US05/27239

ORF3\_19FTW is a cell wall surface protein. An example of an amino acid sequence of ORF3\_19FTW is set forth in SEQ ID NO: 223.

**SEQ ID NO: 223**

5 MKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETS PAIGKVVIKETGEGGALLGDAVFELKNNTDGTTVSQRT  
EAQTGEAIFSNIKPGTYTLTEAOPPVGYKPSTKQWTVEVEKNGRTTVQGEQVENREEALSDQYPQTGTYPDVQTP  
YQIIKVDGSEKNGQHKALNPNPYERVIPEGTLSKRIYQVNNLDDNQYGIELTVSGKTVYERKDKSVPLDVVILLD  
NSNSMSNIRNKNARRAERAGEATRSLIDKITS DPENRVALVTYASTIFDGTEFTVEKGVADKNGKRLNDSLFWNY  
DQTSFTTNTKDYSYLKLTNDKNDIVELKNKVPTEAEDHDGNRLMYQFGATFTQKALMKADEILTQOARQNSQKVI  
10 FHITDGVPTMSYPINFNHATFAPSYQNQLNAFFSKSPNKD GILLSDFITQATSGEHTIVRGDGQSYQMFTDKTVY  
EKGAPAAFPVKPEKYSEMKA VGYAVIGDPINGGYIWLNWRESILAYPFNSNTAKITNHGAPTRWYNGNIAPDGY  
DVFTVGIGINGDPGTDEATATSFMQSISSKPENYTNVTDTTKILEQLNRYFHTIVTEKKSIENTITDPMGELID  
LQLGTDGRFDPADYTLTANDGSRLENGQAVGGPQNDGGLLKNKVFYDTTEKRIRVTGLYLGTGEKVTLTYNVRL  
NDQFVSNKFYDTNGRTTLHPKEVEKNTVRDFPIPKIRDVRKYPAITIAKEKKLGEIEFIKINKNDKKPLRDAVFS  
15 LQKQHPDYPDIYGAI DQNGTYQNVRTGEDGKLT FKNLS DGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRDVT  
SIVPQDIPAGYEFTNDKHYITNEPIPPKREYPRTGGIGMLPFYLGMMMGGVLLYTRKHP

ORF4\_19FTW is a cell wall surface protein. An example of an amino acid sequence of ORF4\_19FTW is set forth in SEQ ID NO: 224.

**SEQ ID NO: 224**

20 MKSINKFLTMLAALLLTASSLFS AATVFAAGTTTT SVTVHKLLATDGDMDKIANELETGNYAGNKVGVL PANAKE  
IAGVMFVWTNTNNEIIDENGQTLGVNIDPQTFKL SGAMPATAMKKLTEAEGAKFNTANLPAKYKIYEIHSLSY  
VGEDGATLTGSKAVPIEIELPLNDVVD AHVYPKNTEAKPKIDKDFKGKANPDTPRVDKDTPVNHQVGDVVEYEIV  
TKIPALANYATANWSDRMTEGLAFNKGTVKVTVD DVALEAGDYALTEVATGFDLKLTDAGLAKVNDQNAEKT VKI  
25 TYSATLNDKAIVEVPESNDVTFNYGNNPDHGNT PKPNKPNENGDLTLTKTWVDATGAPI PAGAEATFDLVNAQTG  
KVVQTVTLTTDKNTVTVNGLDKNT EYKFVERSIKGY SADYQEITTAG EIAVKNWKDENPKPLDPTEPKVV TYGKK  
FVKVNDKDNRLAGAEFVIANADNAGQYLARKADKVS QEEKQLVVTTKDALDRAVAAYNALTAQQQTQQEKEKVDK  
AQAAYNAAVIAANNAFEWVADKDNENVVKLVSDA QGRFEITGLLAGTYYLEETKQPAGYALLTSRQKFEVTATSY  
30 SATGQGI EYTAGSGKDDATKVVNKKITIPQTGGIGTII FAVAGAVIMGI AVYAYVKNNKDEDQLA

ORF5\_19FTW is a cell wall surface protein. An example of an amino acid sequence of ORF5\_19FTW is set forth in SEQ ID NO: 225.

**SEQ ID NO: 225**

35 MTMQKMQMISRIFFVMALCFSLVWGAHAVQAQEDHTLV LQLENYQE VVSQ LPSRDGHR LQVWKLDDSYSDNRV  
QIVRDLHSWDENKLSSFKKT SFEMTFLENQIEVSHIPNGLYYVRSIIQTDAVSYPAEFLFEMTDQTV EPLVIVAK  
KADTVTTKV KLIKVDQDHNRL EGVGFKLVSVARDGSEKEVPLIGEYRYSSSGQVGR TLYTDKNGEIVVTNLPLGT  
YRFKEVEPLAGYTVTTMDTDVQLVDHQLVTITVVNQKLPRGNVDFMKVDGR TNTSLQGAMFKVMKEENGHYTPVL  
QNGKEVVVASGKDGRFRVEGLE YGTYYLWELQAPTGYVQLTSPVSFTIGK DTRKELVTVVKNNKRPRIDVPDTGE  
40 ETLYILMLVAILLFGSGYYLTKKTNN

ORF6\_19FTW is a putative sortase. An example of an amino acid sequence of ORF6\_19FTW is set forth in SEQ ID NO: 226.

**SEQ ID NO: 226**

45 MLIKMAKTKKQKRNNLLLG VVFFIGMAVMAYPLVSRLYYRVESNQ QIADFDKEKATLDEADIDERMKLAQAFNDS  
LNNVSGDPWSEEMKKKGRAEYARMLEIHERM GHVEIPAIDVDLPVYAGTAE EVLQQGAGHLEGTSLPIGGNSTH  
AVITAHTGLPTAKMFTDLTKLKVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
HRLLVRGHRI PYVAEVEEEFIAANKLSHLYRYL FYVAVGLIVILLWIIRRLRKKKRQSERALKALKEATKEVKVE  
DE

ORF7\_19FTW is a putative sortase. An example of an amino acid sequence of ORF7\_19FTW is set forth in SEQ ID NO: 227.

**SEQ ID NO: 227**

MSKSRYSRKKS VKKKKNPFILLI FLVGLAVAMYPLVS RYYYRIESNEVIKEFDET V SQMDKAELEERWRLAQAF  
NATLKPSEILD PFTDQEKKG VSEYANMLKVHERIGYVEI PAIEQEIPMYVGTSEDI LQKGAGLLEGASLPVGGE



NPHTVITAFHRLGLTAELTSQIDKMKKGDTFFLHVLDQVLAYQVDQIVTVEPNDPEPVLIOHGQDYATLLTCTPYM  
INSHRLLVRGKRIPYTAPIAERNRAVRERGGQFWLWLLLGAMAVILLLLYRVYRNRRIVKGLEKQLEGRHVKD

ORF8\_19FTW is a putative sortase. An example of an amino acid sequence of

5 ORF8\_19FTW is set forth in SEQ ID NO: 228.

**SEQ ID NO: 228**

MSRTKLRALLGYLLMLVACLIPIYCFGQMVLSLQSLGQVKGHATFVKSMTTTEMYQEQQNHSLAYNQRLASQNRIVDP  
FLAEGYEVNYQVSDDPDAVYGYLSIPSLEIMEPVYLGADYHHGLMGLAHVDGTPLPLDGTGIRSVIAGHRAEPSH  
VFFRHLDQLKVGDALYYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
10 YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAFGLGILFVLWKLARLLRGK

As discussed above, a *S. pneumoniae* AI sequence is present in the 23F Taiwan 15 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences from 23F Taiwan 15 are set forth below.

15 ORF2\_23FTW is a transcriptional regulator. An example of an amino acid sequence of ORF2\_23FTW is set forth in SEQ ID NO: 229.

**SEQ ID NO: 229**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDTQQVQLIEHHSHQ  
TNYFFHQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISIAATGYRVRQKCGLLLRVGLDLVKNQVVGPEYRIRF  
20 LIALLOFHFGIEIYDLNDGSMDWVTHMIVQSNLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
LKNLFMYPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQLILQHTRGKHLLSKF  
KNILGNDISNLSFLTALTFLTRTFGLQNLVPYNYEYHYGIESDKPLYHISKAIVQEWMTQKIEGVIDQHR  
LYLFSLYLTETIFSSLPAIPIFIILNNQADVNLIKSIILRNFTDKVASVTGYNILISPPPSEEHLEPLIIITTK  
EYLPYVKKQYPKGKHHFLTIALDLHVSQQRLIYQTVDIRKEAFDKRVAMIAKKAHYLL  
25

ORF3\_23FTW is a cell wall surface protein. An example of an amino acid sequence of ORF3\_23FTW is set forth in SEQ ID NO: 230.

**SEQ ID NO: 230**

MKKVRKIFQKAVAGLCCISQLTAFSSIVALAETPETSIPAIGKVVIKETGEGGALLGDAVFELKNNTDGTTVSQRT  
EAQTGEAIFSNIKPGTYTLTEAQQPPVGYKPKSTKQWTVVEVEKNGRRTTVQGEQVENREEALSQYPQTGTYPDVQTP  
YQIIKVDGSEKNGQHKALNPNPYERVIPEGTLISKRIYQVNNLDDNQYGIELTVSGKTVYEQKDKSVPLDVVILLD  
NSNSMSNIRNKNARRAERAGEATRSIDKITSDPENRVALVTYASTIFDGTEFTVEKGVADKNGKRLNDSLFWNY  
DQTSFTTNTKDYSYLKLTNDKNDIVELKNKVPTAEADHDGNRLMYQFGATFTQKALMKADEILTQQARONSQKVI  
FHITDGVPTMSYPINFNHATFAPSQYNQLNAFFSKSPNKDGILLSDFITQATSGEHTIVRGDQSYQMFTDKTVY  
35 EKGAPAAFPVKPEKYSEMKAAGYAVIGDPINGGYIWLNWRESILAYPFNSNTAKITNHGDPTRWYNGNIAPDGY  
DVFTVGIGINGDPGTDEATATSFMQSISSKPENYTNVTDTTKILEQLNRYFHTIVTEKKSIEGNTITDPMGELID  
LQLGTDGRFDPADYTLTANDGSRLENGQAVGGPQNDGGLLKNKVLVDYDTEKRIKRVGLYLGTDEKVTLTYNVRL  
NDEFVSNKFYDTNGRTTLHPKEVEQNTVRDFPIPKIRDVRKYPEITISKEKKLGDIKFNKNDKKPLRDAVFS  
LQKQHPDYPDIYGAIDQNGTYQNVRTGEDGKLTFKNLSDGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRDVT  
40 SIVPQDIPAGYEFTNDKHYITNEPIPPKREYPRGTGGIGMLPFYLGMMMGVLLYTRKHP

ORF4\_23FTW is a cell wall surface protein. An example of an amino acid sequence of ORF4\_23FTW is set forth in SEQ ID NO: 231.

**SEQ ID NO: 231**

MKSINKFLTILAALLLTVSSLSAATVFAAEQKTKTLTVHKLMLTDQELDAWNDAITTAGYDGSQNFQFKQLQ  
GVPQGVTEISGVAFELQSYTGPGQKEQENLTNDVAVTAVNKGVTTETGVKFDTEVLQGTYRLVEVRKESTYVGPN  
GKVLGTGMKAVPALITLPLVNQNGVVENAHVYPKNSEDKPTATKTFDTAAGFVDPGEKGLAIGTKVPYIVTTTIPK  
NSTLATAFWSDEMTEGLDYNQGVVNGQPLDNHYTLEAGHNGFILKLINEKGLEAINGKDAEATITLKYTATL  
NALAVADVPEANDVTFHYGNNGHGNTPKPNKPKNGELTITKTWADAKDAPIAGVEVTFDLVNAQTGEVVKVPGH  
50 ETGIVLNQTNNTFTATGLDNNTEYKFVERTIKGYSADYQTITETGKIAVKNWKDENPEPINPEEPRVKTYGKKF  
VKVDQKDERLKEAQFVVKNEQGKYLALKSAAQAVNEKAAAEAKQALDAAIAAYTNAADKNAAQAVVDAAQKTYN  
DNYRAARFGYVEVERKEDALVLTSTNDGQFQISGLAAGSYTLEETKAPEGFAKLGDVKFEVAGAGSWNQGFNYLK  
DVQKNDATKVVNKKITIPQTGGIGITIIFAVAGAVIMGIAVYAYVKNKDEQDLA



ORF5\_23FTW is a cell wall surface protein. An example of an amino acid sequence of ORF5\_23FTW is set forth in SEQ ID NO: 232.

**SEQ ID NO: 232**

MTMQKMQMISRIFFVMALCFSLVWGAHAVQAQEDHTLVQLQENYQEVVSQLP SRDGHRLQVWKLDDSYSDNRV  
 QIVRDLHSWDENKLSSFKKTSFEMTFLENQIEVSHIPNGLYYVRSIIQTDVASYPAEFLFEMTDQTVEPLVIVAK  
 KADTVTTKVKLIKVDQDHNRLLEGVGFKLVSVARDGSEKEVPLIGEYRYSSSGQVGRITLYTDKNGEIVVTNLPLGT  
 YRKEVEPLAGYTVTTMDTDVQLVDHQLVTITVVNQKLPRGNVDFMKVDGRNTSLQGAMFKVMKEENGHYTPVL  
 QNGKEVVVASGKDGRFRVEGLEYGTYLWELQAPTGYVQLTSPVSFTTIGKDTRKELVTVKNNKRPRIDVPDTGE  
 ETLYILMLVAILLFGSGYYLTCKTNN

ORF6\_23FTW is a putative sortase. An example of an amino acid sequence of ORF6\_23FTW is set forth in SEQ ID NO: 233.

**SEQ ID NO: 233**

MLIKMVTKKKQKRNNLLLGVVFFIGMAVMAYPLVSRLYYRVESNQIADFDKEKATLDEADIDERMKLAQAFNDS  
 LNNVVSGBPSEEMKKKGRAEYARMLEIHERMGHVEIPVIDVDLPVYAGTAEVVLQGGAGQLEGTSPLPIGGNSTH  
 AVITAHTGLPTAKMFTDLTKLVGDKFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
 HRLVLRGHRIPYVAEVEEEFIAANKLSHLRYLRYFVAVGLIVILLWIIRRLRKKKKQPEKALKALKAAARKEVKVE  
 DGQQ

ORF7\_23FTW is a putative sortase. An example of an amino acid sequence of ORF7\_23FTW is set forth in SEQ ID NO: 234.

**SEQ ID NO: 234**

MDNSRRSRKKGTKKKKHPLILLIIFLVGFAVAIYPLVSRYYYRIESNEVIKEFDETVSQMDKAELEERWRLAQAF  
 NATLKPSEILDPEFTEQEKKGVSSEYANMLKVHERIGYVEIPAIQEIIPMYVGTSEEILQKGAGLLEGASLPVGGE  
 NTHTVVTAHRGLPTAELFSQLDKMKKGDVFYLHVLDQVLAYQVDQILTVEPNDFEPVLIQHGKDYATLLTCTPYM  
 INSHRLLVRGKRIPYTAPIAERNRAVRERGQFWLWLLAALVMILVLSYGVYRHRIRIVKGLEKQLEEHVKG

ORF8\_23FTW is a putative sortase. An example of an amino acid sequence of ORF8\_23FTW is set forth in SEQ ID NO: 235.

**SEQ ID NO: 235**

MSKAKLQKLLGYLLMLVALVIPVYCFGQMVLSLQSLGQVKGHEIFSESVTADSYQEQLQRLSDYNQRLDSQNRIVDP  
 FLAEGYEVNYQVSDDPDAVYGYLSIPSLEIMEPVYLGADYHLLAMGLAHVDGTPLPVEGKGIRSVIAGHRAEPSH  
 VFFRHLQDLKVGDALYYDNGQEIYEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
 YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAFGLFVLWKLARLLRGK

As discussed above, a *S. pneumoniae* AI sequence is present in the 23F Poland 16 *S. pneumoniae* genome. Examples of *S. pneumoniae* AI sequences from 23F Poland 16 are set forth below.

ORF2\_23FP is a transcriptional regulator. An example of an amino acid sequence of ORF2\_23FP is set forth in SEQ ID NO: 236.

**SEQ ID NO: 236**

MLNKYIEKRITDKITILNILLDIRSIELDELSTLTSLQSKSLLSILQELQETFEELTFNLDTQQVQLIEHHSQ  
 TNYFHFQLYNQSTILKILRFFLLQGNQSFNEFTQKEYISIATGYRVRQKCGLLLRVGLDLVKNQVVGPEYRIRF  
 LIALLOFHFGEIYDLNDGSMWVTHMIVQNSQLSHELLEITPDEYVHFSILVALTWKRREFPLEFPESKEFEK  
 LKNLFMYPILMEHCQTYLEPHANMTFTQEELDYIFLVYCSANSSFSKDKWNQEKKTHTIQLILQHTRGKHLSSKF  
 KNILGNDISNSLSFLTALTFLTRTFLGLQNLVPYYNYEYHYGIESDKPLYHISKAIQVQEWMTQKIEGVIDQHR  
 LYLFSLYLTETIFSSLPAIPIFIILNNQADVNLKSIILRNFTDKVASVTGYNILISPPPSEEHLTEPLIIITTK  
 EYLPYVKKQYPKGKHHFLTIALDLHVSQQRLIYQTIIVDIRKEAFDKRVAMIAKKAHYLL

ORF3\_23FP is a cell wall surface protein. An example of an amino acid sequence of ORF3\_23FP is set forth in SEQ ID NO: 237.

**SEQ ID NO: 237**



MKKVRKLEQKAVATLCTISQUTAFSSIVALLAETPETSIPAIGKVVIKETGEGGALLGDAVFELKNNTDGTTVSQRTEAQTGEAIFSNIKPGTYTLTEAQPPVGYKPKSTKQWTVEVEKNGRRTTVQGEQVENREEALSDQYPQTGTYPDVQTPYQIIKVDGSEKNGQHKALNPNPYERVIPEGTLISKRIYQVNNLDDNQYGIELTVSGKTTVETKEASTPLDVVILLDNSNSMSNIRHNHAHRAEKAGEATRALVDKITSNPDNRVALVTYGSTIFDGSEATVEKGVADANGKILNDSALWTF  
 5 DRTTFTAKTYNYSFLNLTSDPTDIQTIKDRIPSDAEELNKDKLAMYQFGATFTQKALMTADDILTKQARPNSKKVIFHITDGVPTMSYPINFKYTGTTQSYRTQLNNFKAKTPNSSGILLEDFVTWSADGEHKIVRGDGESYQMFTKKPVT  
 DQYGVHQILSITSMEQRAKLVSAGYRFYGTDLVLYWRDSILAYPFNSSTDWITNHGDPPTWYNGNMAQDGYDVF  
 TVGVGVNGDPGTDEATATRFMQSISSSPDNYTNVADPSQILQELNRYFYTIVNEKKSIENTITDPMGELIDFQLGADGRFDPADYTLTANDGSSLVNNVPTGGPQNDGGLLKNKAVFYDTTEKRIRVTGLYLGTGEKVTLTYNVRLNDQ  
 10 FVSNKFYDTNGRTTLHPKEVEKNTVRDFPIPKIRDVRKYPEITIPKEKKLGEIEFIKINKNDKKPLRDAVFSLOKQHPDYFDIYGAIDQNGTYQNVRTGEDGKLTFKNLSDGKYRLFENSEPAGYKPVQNKPIVAFQIVNGEVRDVTISIV  
 PQDIPAGYEFTNDKHYITNEPIPPKREYPRRTGGIGMLPFYLGCMMSGGVLLYTRKNP

ORF4\_23FP is a cell wall surface protein. An example of an amino acid sequence of  
 ORF4\_23FP is set forth in SEQ ID NO: 238.

15 **SEQ ID NO: 238**

MKSINKFLTMLAALLLTASSLFSAA TVFAADNVSTAPDAVTKTLTIHKLLLSDDLKTWDTNGPKGYDGTQSSSLK  
 DLTGVVAEEIPNVYFELQKYNLT DGKEKENLKDDSKWTTVHGGLTTKDGLKIETSTLKGVYRIREDRTKTTYVGP  
 NGQVLTGSKAVPALVTLPLVNNNGTVIDAHVF PKNSYNKPVDKRIADTLNNDQNGLSIGTKIPYVVNTTIPSN  
 ATFATSFWSDEMTEGLTYNEDVTITLNNVAMDQADYEVTGKINGFNLKLT EAGLAKINGKDADQKIQITYSATLN  
 20 SLAVADIPESNDITYHYGNHQDHGNTPKPTKPNNGQITVTKTWD SQPAPEGVKATVQLVNAKTGEKVGAPVELSE  
 NNWTYTWSGLDNSIEYKVEEYNGYSAEYTVESKGLGVKNWKDNNPAPINLEEPRVKTYGKKFVKVDQKDRLE  
 NAQFVVKKADSNKYIAFKSTAQQAADEKAAATAKQKLDAAVAAYTNAADKQAAQALVDQAQQEYNVAYKEAKFGY  
 VEVAGKDEAMVLTSNTDGQFQISGLAAGTYKLEEIKAPEGFAKIDDEFFVVGAGSWNQGEFNYLKDVQKNDATKV  
 VNKKITIPQTGGIGTIIFAVAGAVIMGIAVYAYVKNKDEDDQLA

25 ORF5\_23FP is a cell wall surface protein. An example of an amino acid sequence of  
 ORF5\_23FP is set forth in SEQ ID NO: 239.

**SEQ ID NO: 239**

MTMQKMQMISRIFFVMALCFSLVWGAHAVQAQEDHTLVLQLENYQE VVSQ LPSRDGHR LQVWKLDDSYSDNRV  
 30 QIVRDLHSWDENKLSSFKKT SFEMTFLENQIEVSHIPNGLYYVRSIIQTD AVSYPAEFLFEMTDQTV EPLVIVAK  
 KADTVTTKV KLIKVDQDHNRL EGVGFKLVSVARDGSEKEVPLIGEYRYSSSGQVGR TLYTDKNGEI VVTNLPLGT  
 YRFKEVEPLAGYAVTTMDTDVQLVDHQLVTITVNVQKLPRGNVDFMKVDGR TNTSLQGAMFKVMKEENGHYTPVL  
 QNGKEVVVASGKDGFRVEGLE YGTYYLWELQAPTGYVQLTSPVSFTIGKDTRKELVT VVKNNKRPRIDVPDTGE  
 35 ETLYILMLVAILLFGSGYYLT KKTNN

ORF6\_23FP is a putative sortase. An example of an amino acid sequence of ORF6\_23FP is  
 set forth in SEQ ID NO: 240.

**SEQ ID NO: 240**

40 MLIKMAKTKKQKRNNLLLG VVFFIGIAVMAYPLVSRLYYRVESNQ QIADFDKEKATLDEADI DERMKLAQAFNDS  
 LNNVVS GDPWSEEMKKKGRAEYARMLEIHERM GHVEIPAI DVDLPVYAGTAE EVLQQGAGHLEGTSLPIGGNSTH  
 AVITAHTGLPTAKMFTDLTKLKVGD KFYVHNIKEVMAYQVDQVKVIEPTNFDDLLIVPGHDYVTLTCTPYMINT  
 HRLLRGHRIPYVAEVEEEFIAANKLSHLYRYLFYVAVGLIVILLWII RRLRKKKRQSERALKALKEATKEVKVE  
 DE

45 ORF7\_23FP is a putative sortase. An example of an amino acid sequence of ORF7\_23FP is  
 set forth in SEQ ID NO: 241.

**SEQ ID NO: 241**

50 MSKSRYSRKKS VKKKNPFILLIIFLVGLAVAMYPLVSRYYYRIESNEVIKEFDET V SQMDKAELEERWRLAQAF  
 NATLKPSEILDPFTEQEKKKG VSEYANMLKVHERIGYVEIPAI DQEI PMYVGTSEEILQKGAGLLEGASLPVGGE  
 NTHTVVTAHRGLPTAE LFSQLDKMKKGDI FYLHVLDQVLAYQVDQIVTVEPNDFEPVLIQHGEDYATLLTCTPYM  
 INSHRLVRGKRIPYTAPIAERNRAVRERGQFWLWLLLGAMAVILL LLYRVYRNRRIVKGLEKQLEGRHVKD

ORF8\_23FP is a putative sortase. An example of an amino acid sequence of ORF8\_23FP is  
 set forth in SEQ ID NO: 242.



SEQ ID NO: 242

MSRTKLRALLGYLLMLVACLIPIYCFGQMVLSLQGVKGHATFVKSMTTTEMYQEQQNHSLAYNQRLASQNRIVDP  
FLAEGYEVNYQVSDDDPAVYGYLSIPSLEIMEPVYLGADYHHLGMGLAHVDGTPLPLDGTGIRSVIAGHRAEPSH  
VFFRHLDLQKVGDALYYDNGQEIVEYQMMDEIILPSEWEKLESVSSKNIMTLITCDPIPTFNKRLLVNFERVAV  
YQKSDPQTAAVARVAFTKEGQSVSRVATSQWLYRGLVVLAFGLFVLWKLARLLRGK

Immunogenic compositions of the invention comprising AI antigens may further comprise one or more antigenic agents. Preferred antigens include those listed below. Additionally, the compositions of the present invention may be used to treat or prevent infections caused by any of the below-listed microbes. Antigens for use in the immunogenic compositions include, but are not limited to, one or more of the following set forth below, or antigens derived from one or more of the following set forth below:

Bacterial Antigens

*N. meningitides*: a protein antigen from *N. meningitides* serogroup A, C, W135, Y, and/or B (1-7); an outer-membrane vesicle (OMV) preparation from *N. meningitides* serogroup B. (8, 9, 10, 11); a saccharide antigen, including LPS, from *N. meningitides* serogroup A, B, C W135 and/or Y, such as the oligosaccharide from serogroup C (see PCT/US99/09346; PCT IB98/01665; and PCT IB99/00103);

*Streptococcus pneumoniae*: a saccharide or protein antigen, particularly a saccharide from *Streptococcus pneumoniae*;

*Streptococcus agalactiae*: particularly, Group B streptococcus antigens;

*Streptococcus pyogenes*: particularly, Group A streptococcus antigens;

*Enterococcus faecalis* or *Enterococcus faecium*: Particularly a trisaccharide repeat or other *Enterococcus* derived antigens provided in US Patent No. 6,756,361;

*Helicobacter pylori*: including: Cag, Vac, Nap, HopX, HopY and/or urease antigen;

*Bordetella pertussis*: such as pertussis holotoxin (PT) and filamentous haemagglutinin (FHA) from *B. pertussis*, optionally also combination with pertactin and/or agglutinogens 2 and 3 antigen;

*Staphylococcus aureus*: including *S. aureus* type 5 and 8 capsular polysaccharides optionally conjugated to nontoxic recombinant *Pseudomonas aeruginosa* exotoxin A, such as StaphVAX™, or antigens derived from surface proteins, invasins (leukocidin, kinases, hyaluronidase), surface factors that inhibit phagocytic engulfment (capsule, Protein A), carotenoids, catalase production, Protein A, coagulase, clotting factor, and/or membrane-damaging toxins (optionally detoxified) that lyse eukaryotic cell membranes (hemolysins, leukotoxin, leukocidin);

*Staphylococcus epidermis*: particularly, *S. epidermidis* slime-associated antigen (SAA);

*Staphylococcus saprophyticus*: (causing urinary tract infections) particularly the 160 kDa hemagglutinin of *S. saprophyticus* antigen;

*Pseudomonas aeruginosa*: particularly, endotoxin A, Wzz protein, *P. aeruginosa* LPS, more particularly LPS isolated from PAO1 (O5 serotype), and/or Outer Membrane Proteins, including Outer Membrane Proteins F (OprF) (*Infect Immun.* 2001 May; 69(5): 3510-3515);



~~of the *Bacillus anthracis* (anthrax) such as *B. anthracis* antigens (optionally detoxified) from A-components (lethal factor (LF) and edema factor (EF)), both of which can share a common B-component known as protective antigen (PA);~~

*Moraxella catarrhalis*: (respiratory) including outer membrane protein antigens (HMW-OMP), C-antigen, and/or LPS;

*Yersinia pestis* (plague): such as F1 capsular antigen (*Infect Immun.* 2003 Jan; 71(1): 374-383, LPS (*Infect Immun.* 1999 Oct; 67(10): 5395), *Yersinia pestis* V antigen (*Infect Immun.* 1997 Nov; 65(11): 4476-4482);

*Yersinia enterocolitica* (gastrointestinal pathogen): particularly LPS (*Infect Immun.* 2002 August; 70(8): 4414);

*Yersinia pseudotuberculosis*: gastrointestinal pathogen antigens;

*Mycobacterium tuberculosis*: such as lipoproteins, LPS, BCG antigens, a fusion protein of antigen 85B (Ag85B) and/or ESAT-6 optionally formulated in cationic lipid vesicles (*Infect Immun.* 2004 October; 72(10): 6148), *Mycobacterium tuberculosis* (Mtb) isocitrate dehydrogenase associated antigens (*Proc Natl Acad Sci U S A.* 2004 Aug 24; 101(34): 12652), and/or MPT51 antigens (*Infect Immun.* 2004 July; 72(7): 3829);

*Legionella pneumophila* (Legionnaires' Disease): *L. pneumophila* antigens -- optionally derived from cell lines with disrupted *asd* genes (*Infect Immun.* 1998 May; 66(5): 1898);

*Rickettsia*: including outer membrane proteins, including the outer membrane protein A and/or B (OmpB) (*Biochim Biophys Acta.* 2004 Nov 1; 1702(2):145), LPS, and surface protein antigen (SPA) (*J Autoimmun.* 1989 Jun; 2 Suppl:81);

*E. coli*: including antigens from enterotoxigenic *E. coli* (ETEC), enteroaggregative *E. coli* (EAaggEC), diffusely adhering *E. coli* (DAEC), enteropathogenic *E. coli* (EPEC), and/or enterohemorrhagic *E. coli* (EHEC);

*Vibrio cholerae*: including proteinase antigens, LPS, particularly lipopolysaccharides of *Vibrio cholerae* II, O1 Inaba O-specific polysaccharides, *V. cholera* O139, antigens of IEM108 vaccine (*Infect Immun.* 2003 Oct; 71(10):5498-504), and/or Zonula occludens toxin (Zot);

*Salmonella typhi* (typhoid fever): including capsular polysaccharides preferably conjugates (Vi, i.e. vax-TyVi);

*Salmonella typhimurium* (gastroenteritis): antigens derived therefrom are contemplated for microbial and cancer therapies, including angiogenesis inhibition and modulation of flk;

*Listeria monocytogenes* (systemic infections in immunocompromised or elderly people, infections of fetus): antigens derived from *L. monocytogenes* are preferably used as carriers/vectors for intracytoplasmic delivery of conjugates/associated compositions of the present invention;

*Porphyromonas gingivalis*: particularly, *P. gingivalis* outer membrane protein (OMP);

*Tetanus*: such as tetanus toxoid (TT) antigens, preferably used as a carrier protein in conjunction/conjugated with the compositions of the present invention;

~~PCT Diphtheria~~ such as a diphtheria toxoid, preferably CRM<sub>197</sub>, additionally antigens capable of modulating, inhibiting or associated with ADP ribosylation are contemplated for combination/co-administration/conjugation with the compositions of the present invention, the diphtheria toxoids are preferably used as carrier proteins;

5 *Borrelia burgdorferi* (Lyme disease): such as antigens associated with P39 and P13 (an integral membrane protein, *Infect Immun.* 2001 May; 69(5): 3323-3334), VlsE Antigenic Variation Protein (*J Clin Microbiol.* 1999 Dec; 37(12): 3997);

*Haemophilus influenzae B*: such as a saccharide antigen therefrom;

10 *Klebsiella*: such as an OMP, including OMP A, or a polysaccharide optionally conjugated to tetanus toxoid;

*Neisseria gonorrhoeae*: including, a Por (or porin) protein, such as PorB (*see Zhu et al., Vaccine* (2004) 22:660 – 669), a transferring binding protein, such as TbpA and TbpB (*See Price et al., Infection and Immunity* (2004) 71(1):277 – 283), a opacity protein (such as Opa), a reduction-modifiable protein (Rmp), and outer membrane vesicle (OMV) preparations (*see Plante et al., J Infectious Disease* (2000) 182:848 – 855), also see *e.g.* WO99/24578, WO99/36544, WO99/57280, WO02/079243);

*Chlamydia pneumoniae*: particularly *C. pneumoniae* protein antigens;

20 *Chlamydia trachomatis*: including antigens derived from serotypes A, B, Ba and C are (agents of trachoma, a cause of blindness), serotypes L<sub>1</sub>, L<sub>2</sub> & L<sub>3</sub> (associated with Lymphogranuloma venereum), and serotypes, D-K;

*Treponema pallidum* (Syphilis): particularly a TmpA antigen; and

*Haemophilus ducreyi* (causing chancroid): including outer membrane protein (DsrA).

25 Where not specifically referenced, further bacterial antigens of the invention may be capsular antigens, polysaccharide antigens or protein antigens of any of the above. Further bacterial antigens may also include an outer membrane vesicle (OMV) preparation. Additionally, antigens include live, attenuated, split, and/or purified versions of any of the aforementioned bacteria. The bacterial or microbial derived antigens of the present invention may be gram-negative or gram-positive and aerobic or anaerobic.

30 Additionally, any of the above bacterial-derived saccharides (polysaccharides, LPS, LOS or oligosaccharides) can be conjugated to another agent or antigen, such as a carrier protein (for example CRM<sub>197</sub>). Such conjugation may be direct conjugation effected by reductive amination of carbonyl moieties on the saccharide to amino groups on the protein, as provided in US Patent No. 5,360,897 and *Can J Biochem Cell Biol.* 1984 May;62(5):270-5. Alternatively, the saccharides can be conjugated through a linker, such as, with succinamide or other linkages provided in *Bioconjugate Techniques*, 1996 and *CRC, Chemistry of Protein Conjugation and Cross-Linking*, 1993.



Viral Antigens

*Influenza*: including whole viral particles (attenuated), split, or subunit comprising hemagglutinin (HA) and/or neuraminidase (NA) surface proteins, the influenza antigens may be derived from chicken embryos or propagated on cell culture, and/or the influenza antigens are derived from influenza type A, B, and/or C, among others;

*Respiratory syncytial virus (RSV)*: including the F protein of the A2 strain of RSV (*J Gen Virol.* 2004 Nov; 85(Pt 11):3229) and/or G glycoprotein;

*Parainfluenza virus (PIV)*: including PIV type 1, 2, and 3, preferably containing hemagglutinin, neuraminidase and/or fusion glycoproteins;

*Poliovirus*: including antigens from a family of picornaviridae, preferably poliovirus antigens such as OPV or, preferably IPV;

*Measles*: including split measles virus (MV) antigen optionally combined with the Protollin and or antigens present in MMR vaccine;

*Mumps*: including antigens present in MMR vaccine;

*Rubella*: including antigens present in MMR vaccine as well as other antigens from Togaviridae, including dengue virus;

*Rabies*: such as lyophilized inactivated virus (RabAvert™);

*Flaviviridae viruses*: such as (and antigens derived therefrom) yellow fever virus, Japanese encephalitis virus, dengue virus (types 1, 2, 3, or 4), tick borne encephalitis virus, and West Nile virus;

*Caliciviridae*; antigens therefrom;

*HIV*: including HIV-1 or HIV-2 strain antigens, such as gag (p24gag and p55gag), env (gp160 and gp41), pol, tat, nef, rev vpu, miniproteins, (preferably p55 gag and gp140v delete) and antigens from the isolates HIV<sub>IIIB</sub>, HIV<sub>SF2</sub>, HIV<sub>LAV</sub>, HIV<sub>LAI</sub>, HIV<sub>MN</sub>, HIV-1<sub>CM235</sub>, HIV-1<sub>US4</sub>, HIV-2; simian immunodeficiency virus (SIV) among others;

*Rotavirus*: including VP4, VP5, VP6, VP7, VP8 proteins (*Protein Expr Purif.* 2004 Dec;38(2):205) and/or NSP4;

*Pestivirus*: such as antigens from classical porcine fever virus, bovine viral diarrhoea virus, and/or border disease virus;

*Parvovirus*: such as parvovirus B19;

*Coronavirus*: including SARS virus antigens, particularly spike protein or proteases therefrom, as well as antigens included in WO 04/92360;

*Hepatitis A virus*: such as inactivated virus;

*Hepatitis B virus*: such as the surface and/or core antigens (sAg), as well as the presurface sequences, pre-S1 and pre-S2 (formerly called pre-S), as well as combinations of the above, such as sAg/pre-S1, sAg/pre-S2, sAg/pre-S1/pre-S2, and pre-S1/pre-S2, (see, e.g., AHBV Vaccines - *Human Vaccines and Vaccination*, pp. 159-176; and U.S. Patent Nos. 4,722,840, 5,098,704, 5,324,513;

Beck et al., *J. Virol.* (1995) 69:6833-6838, Birnbaum et al., *J. Virol.* (1990) 64:3319-3330; and Zhou et al., *J. Virol.* (1991) 65:5457-5464);

*Hepatitis C virus*: such as E1, E2, E1/E2 (see, Houghton et al., *Hepatology* (1991) 14:381), NS345 polyprotein, NS 345-core polyprotein, core, and/or peptides from the nonstructural regions (International Publication Nos. WO 89/04669; WO 90/11089; and WO 90/14436);

*Delta hepatitis virus (HDV)*: antigens derived therefrom, particularly  $\delta$ -antigen from HDV (see, e.g., U.S. Patent No. 5,378,814);

*Hepatitis E virus (HEV)*; antigens derived therefrom;

*Hepatitis G virus (HGV)*; antigens derived therefrom;

*Varicella zoster virus*: antigens derived from varicella zoster virus (VZV) (*J. Gen. Virol.* (1986) 67:1759);

*Epstein-Barr virus*: antigens derived from EBV (Baer et al., *Nature* (1984) 310:207);

*Cytomegalovirus*: CMV antigens, including gB and gH (*Cytomegaloviruses* (J.K. McDougall, ed., Springer-Verlag 1990) pp. 125-169);

*Herpes simplex virus*: including antigens from HSV-1 or HSV-2 strains and glycoproteins gB, gD and gH (McGeoch et al., *J. Gen. Virol.* (1988) 69:1531 and U.S. Patent No. 5,171,568);

*Human Herpes Virus*: antigens derived from other human herpesviruses such as HHV6 and HHV7; and

*HPV*: including antigens associated with or derived from human papillomavirus (HPV), for example, one or more of E1 – E7, L1, L2, and fusions thereof, particularly the compositions of the invention may include a virus-like particle (VLP) comprising the L1 major capsid protein, more particular still, the HPV antigens are protective against one or more of HPV serotypes 6, 11, 16 and/or 18.

Further provided are antigens, compositions, methods, and microbes included in *Vaccines*, 4<sup>th</sup> Edition (Plotkin and Orenstein ed. 2004); *Medical Microbiology* 4<sup>th</sup> Edition (Murray et al. ed. 2002); *Virology*, 3rd Edition (W.K. Joklik ed. 1988); *Fundamental Virology*, 2nd Edition (B.N. Fields and D.M. Knipe, eds. 1991), which are contemplated in conjunction with the compositions of the present invention.

Additionally, antigens include live, attenuated, split, and/or purified versions of any of the aforementioned viruses.

#### Fungal Antigens

Fungal antigens for use herein, associated with vaccines include those described in: U.S. Pat. Nos. 4,229,434 and 4,368,191 for prophylaxis and treatment of trichophytosis caused by Trichophyton mentagrophytes; U.S. Pat. Nos. 5,277,904 and 5,284,652 for a broad spectrum dermatophyte vaccine for the prophylaxis of dermatophyte infection in animals, such as guinea pigs, cats, rabbits, horses and lambs, these antigens comprises a suspension of killed *T. equinum*, *T. mentagrophytes* (var. granulare), *M. canis* and/or *M. gypseum* in an effective amount optionally combined with an adjuvant;



U.S. Pat. Nos. 5,453,273 and 6,132,733 for a ringworm vaccine comprising an effective amount of a homogenized, formaldehyde-killed fungi, i.e., *Microsporum canis* culture in a carrier; U.S. Pat. No. 5,948,413 involving extracellular and intracellular proteins for pythiosis. Additional antigens identified within antifungal vaccines include Ringvac bovis LTF-130 and Bioveta.

Further, fungal antigens for use herein may be derived from Dermatophytes, including: *Epidermophyton floccusum*, *Microsporum audouini*, *Microsporum canis*, *Microsporum distortum*, *Microsporum equinum*, *Microsporum gypsum*, *Microsporum nanum*, *Trichophyton concentricum*, *Trichophyton equinum*, *Trichophyton gallinae*, *Trichophyton gypseum*, *Trichophyton megnini*, *Trichophyton mentagrophytes*, *Trichophyton quinckeanum*, *Trichophyton rubrum*, *Trichophyton schoenleini*, *Trichophyton tonsurans*, *Trichophyton verrucosum*, *T. verrucosum* var. album, var. discoides, var. ochraceum, *Trichophyton violaceum*, and/or *Trichophyton faviforme*.

Fungal pathogens for use as antigens or in derivation of antigens in conjunction with the compositions of the present invention comprise *Aspergillus fumigatus*, *Aspergillus flavus*, *Aspergillus niger*, *Aspergillus nidulans*, *Aspergillus terreus*, *Aspergillus sydowi*, *Aspergillus flavatus*, *Aspergillus glaucus*, *Blastoschizomyces capitatus*, *Candida albicans*, *Candida enolase*, *Candida tropicalis*, *Candida glabrata*, *Candida krusei*, *Candida parapsilosis*, *Candida stellatoidea*, *Candida kusei*, *Candida parakwsei*, *Candida lusitaniae*, *Candida pseudotropicalis*, *Candida guilliermondi*, *Cladosporium carrionii*, *Coccidioides immitis*, *Blastomyces dermatidis*, *Cryptococcus neoformans*, *Geotrichum clavatum*, *Histoplasma capsulatum*, *Klebsiella pneumoniae*, *Paracoccidioides brasiliensis*, *Pneumocystis carinii*, *Pythium insidiosum*, *Pityrosporum ovale*, *Sacharomyces cerevisiae*, *Saccharomyces boulardii*, *Saccharomyces pombe*, *Scedosporium apiospermum*, *Sporothrix schenckii*, *Trichosporon beigeli*, *Toxoplasma gondii*, *Penicillium marneffe*, *Malassezia* spp., *Fonsecaea* spp., *Wangiella* spp., *Sporothrix* spp., *Basidiobolus* spp., *Conidiobolus* spp., *Rhizopus* spp., *Mucor* spp., *Absidia* spp., *Mortierella* spp., *Cunninghamella* spp., and *Saksenaea* spp.

Other fungi from which antigens are derived include *Alternaria* spp., *Curvularia* spp., *Helminthosporium* spp., *Fusarium* spp., *Aspergillus* spp., *Penicillium* spp., *Monolinia* spp., *Rhizoctonia* spp., *Paecilomyces* spp., *Pithomyces* spp., and *Cladosporium* spp.

Processes for producing a fungal antigens are well known in the art (see US Patent No. 6,333,164). In a preferred method a solubilized fraction extracted and separated from an insoluble fraction obtainable from fungal cells of which cell wall has been substantially removed or at least partially removed, characterized in that the process comprises the steps of: obtaining living fungal cells; obtaining fungal cells of which cell wall has been substantially removed or at least partially removed; bursting the fungal cells of which cell wall has been substantially removed or at least partially removed; obtaining an insoluble fraction; and extracting and separating a solubilized fraction from the insoluble fraction.

STD Antigens

In particular embodiments, microbes (bacteria, viruses and/or fungi) against which the present compositions and methods can be implemented include those that cause sexually transmitted diseases (STDs) and/or those that display on their surface an antigen that can be the target or antigen composition of the invention. In a preferred embodiment of the invention, compositions are combined with antigens derived from a viral or bacterial STD. Antigens derived from bacteria or viruses can be administered in conjunction with the compositions of the present invention to provide protection against at least one of the following STDs, among others: chlamydia, genital herpes, hepatitis (particularly HCV), genital warts, gonorrhoea, syphilis and/or chancroid (See, WO00/15255).

In another embodiment the compositions of the present invention are co-administered with an antigen for the prevention or treatment of an STD.

Antigens derived from the following viruses associated with STDs, which are described in greater detail above, are preferred for co-administration with the compositions of the present invention: hepatitis (particularly HCV), HPV, HIV, or HSV.

Additionally, antigens derived from the following bacteria associated with STDs, which are described in greater detail above, are preferred for co-administration with the compositions of the present invention: *Neisseria gonorrhoeae*, *Chlamydia pneumoniae*, *Chlamydia trachomatis*, *Treponema pallidum*, or *Haemophilus ducreyi*.

Respiratory Antigens

The antigen may be a respiratory antigen and could further be used in an immunogenic composition for methods of preventing and/or treating infection by a respiratory pathogen, including a virus, bacteria, or fungi such as respiratory syncytial virus (RSV), PIV, SARS virus, influenza, *Bacillus anthracis*, particularly by reducing or preventing infection and/or one or more symptoms of respiratory virus infection. A composition comprising an antigen described herein, such as one derived from a respiratory virus, bacteria or fungus is administered in conjunction with the compositions of the present invention to an individual which is at risk of being exposed to that particular respiratory microbe, has been exposed to a respiratory microbe or is infected with a respiratory virus, bacteria or fungus. The composition(s) of the present invention is/are preferably co-administered at the same time or in the same formulation with an antigen of the respiratory pathogen. Administration of the composition results in reduced incidence and/or severity of one or more symptoms of respiratory infection.

Pediatric/Geriatric Antigens

In one embodiment the compositions of the present invention are used in conjunction with an antigen for treatment of a pediatric population, as in a pediatric antigen. In a more particular embodiment the pediatric population is less than about 3 years old, or less than about 2 years, or less than about 1 years old. In another embodiment the pediatric antigen (in conjunction with the composition of the present invention) is administered multiple times over at least 1, 2, or 3 years.



In another embodiment, the compositions of the present invention are used in conjunction with an antigen for treatment of a geriatric population, as in a geriatric antigen.

#### Other Antigens

Other antigens for use in conjunction with the compositions of the present include hospital acquired (nosocomial) associated antigens.

In another embodiment, parasitic antigens are contemplated in conjunction with the compositions of the present invention. Examples of parasitic antigens include those derived from organisms causing malaria and/or Lyme disease.

In another embodiment, the antigens in conjunction with the compositions of the present invention are associated with or effective against a mosquito born illness. In another embodiment, the antigens in conjunction with the compositions of the present invention are associated with or effective against encephalitis. In another embodiment the antigens in conjunction with the compositions of the present invention are associated with or effective against an infection of the nervous system.

In another embodiment, the antigens in conjunction with the compositions of the present invention are antigens transmissible through blood or body fluids.

#### Antigen Formulations

In other aspects of the invention, methods of producing microparticles having adsorbed antigens are provided. The methods comprise: (a) providing an emulsion by dispersing a mixture comprising (i) water, (ii) a detergent, (iii) an organic solvent, and (iv) a biodegradable polymer selected from the group consisting of a poly( $\alpha$ -hydroxy acid), a polyhydroxy butyric acid, a polycaprolactone, a polyorthoester, a polyanhydride, and a polycyanoacrylate. The polymer is typically present in the mixture at a concentration of about 1% to about 30% relative to the organic solvent, while the detergent is typically present in the mixture at a weight-to-weight detergent-to-polymer ratio of from about 0.00001:1 to about 0.1:1 (more typically about 0.0001:1 to about 0.1:1, about 0.001:1 to about 0.1:1, or about 0.005:1 to about 0.1:1); (b) removing the organic solvent from the emulsion; and (c) adsorbing an antigen on the surface of the microparticles. In certain embodiments, the biodegradable polymer is present at a concentration of about 3% to about 10% relative to the organic solvent.

Microparticles for use herein will be formed from materials that are sterilizable, non-toxic and biodegradable. Such materials include, without limitation, poly( $\alpha$ -hydroxy acid), polyhydroxybutyric acid, polycaprolactone, polyorthoester, polyanhydride, PACA, and polycyanoacrylate. Preferably, microparticles for use with the present invention are derived from a poly( $\alpha$ -hydroxy acid), in particular, from a poly(lactide) ("PLA") or a copolymer of D,L-lactide and glycolide or glycolic acid, such as a poly(D,L-lactide-co-glycolide) ("PLG" or "PLGA"), or a copolymer of D,L-lactide and caprolactone. The microparticles may be derived from any of various polymeric starting materials which have a variety of molecular weights and, in the case of the copolymers such as PLG, a variety of lactide:glycolide ratios, the selection of which will be largely a

matter of choice, depending in part on the coadministered macromolecule. These parameters are discussed more fully below.

Further antigens may also include an outer membrane vesicle (OMV) preparation.

Additional formulation methods and antigens (especially tumor antigens) are provided in U.S.

5 Patent Serial No. 09/581,772.

#### Antigen References

The following references include antigens useful in conjunction with the compositions of the present invention:

- 10 1 International patent application WO99/24578
- 2 International patent application WO99/36544.
- 3 International patent application WO99/57280.
- 4 International patent application WO00/22430.
- 5 Tettelin et al. (2000) Science 287:1809-1815.
- 15 6 International patent application WO96/29412.
- 7 Pizza et al. (2000) Science 287:1816-1820.
- 8 PCT WO 01/52885.
- 9 Bjune et al. (1991) Lancet 338(8775).
- 10 Fuskasawa et al. (1999) Vaccine 17:2951-2958.
- 20 11 Rosenqist et al. (1998) Dev. Biol. Strand 92:323-333.
- 12 Constantino et al. (1992) Vaccine 10:691-698.
- 13 Constantino et al. (1999) Vaccine 17:1251-1263.
- 14 Watson (2000) Pediatr Infect Dis J 19:331-332.
- 15 Rubin (2000) Pediatr Clin North Am 47:269-285,v.
- 25 16 Jedrzejewski (2001) Microbiol Mol Biol Rev 65:187-207.
- 17 International patent application filed on 3<sup>rd</sup> July 2001 claiming priority from GB-0016363.4; WO 02/02606; PCT IB/01/00166.
- 18 Kalman et al. (1999) Nature Genetics 21:385-389.
- 19 Read et al. (2000) Nucleic Acids Res 28:1397-406.
- 30 20 Shirai et al. (2000) J. Infect. Dis 181(Suppl 3):S524-S527.
- 21 International patent application WO99/27105.
- 22 International patent application WO00/27994.
- 23 International patent application WO00/37494.
- 24 International patent application WO99/28475.
- 35 25 Bell (2000) Pediatr Infect Dis J 19:1187-1188.
- 26 Iwarson (1995) APMIS 103:321-326.
- 27 Gerlich et al. (1990) Vaccine 8 Suppl:S63-68 & 79-80.
- 28 Hsu et al. (1999) Clin Liver Dis 3:901-915.
- 29 Gastofsson et al. (1996) N. Engl. J. Med. 334-:349-355.
- 40 30 Rappuoli et al. (1991) TIBTECH 9:232-238.
- 31 Vaccines (1988) eds. Plotkin & Mortimer. ISBN 0-7216-1946-0.
- 32 Del Giudice et al. (1998) Molecular Aspects of Medicine 19:1-70.
- 33 International patent application WO93/018150.
- 34 International patent application WO99/53310.
- 45 35 International patent application WO98/04702.
- 36 Ross et al. (2001) Vaccine 19:135-142.
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15 56 Hermanson (1996) *Bioconjugate Techniques* ISBN: 012323368 & 012342335X.

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59 European patent application 0427347.

60 International patent application WO93/17712.

20 61 International patent application WO98/58668.

62 European patent application 0471177.

63 International patent application WO00/56360.

64 International patent application WO00/67161.

25 The contents of all of the above cited patents, patent applications and journal articles are incorporated by reference as if set forth fully herein.

There may be an upper limit to the number of Gram positive bacterial proteins which will be in the compositions of the invention. Preferably, the number of Gram positive bacterial proteins in a composition of the invention is less than 20, less than 19, less than 18, less than 17, less than 16, less than 15, less than 14, less than 13, less than 12, less than 11, less than 10, less than 9, less than 8, less than 7, less than 6, less than 5, less than 4, or less than 3. Still more preferably, the number of Gram positive bacterial proteins in a composition of the invention is less than 6, less than 5, or less than 4. Still more preferably, the number of Gram positive bacterial proteins in a composition of the invention is 3.

35 The Gram positive bacterial proteins and polynucleotides used in the invention are preferably isolated, *i.e.*, separate and discrete, from the whole organism with which the molecule is found in nature or, when the polynucleotide or polypeptide is not found in nature, is sufficiently free of other biological macromolecules so that the polynucleotide or polypeptide can be used for its intended purpose.

#### 40 Fusion Proteins: GBS AI sequences

The GBS AI proteins used in the invention may be present in the composition as individual separate polypeptides, but it is preferred that at least two (*i.e.* 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, or 18) of the antigens are expressed as a single polypeptide chain (a "hybrid" or "fusion" polypeptide). Such fusion polypeptides offer two principal advantages: first, a polypeptide that may be unstable or poorly expressed on its own can be assisted by adding a suitable fusion partner that

overcomes the problem; second, commercial manufacture is simplified as only one expression and purification need be employed in order to produce two polypeptides which are both antigenically useful.

The fusion polypeptide may comprise one or more AI polypeptide sequences. Preferably, the fusion comprises an AI surface protein sequence. Preferably, the fusion polypeptide includes one or more of GBS 80, GBS 104, and GBS 67. Most preferably, the fusion peptide includes a polypeptide sequence from GBS 80. Accordingly, the invention includes a fusion peptide comprising a first amino acid sequence and a second amino acid sequence, wherein said first and second amino acid sequences are selected from a GBS AI surface protein or a fragment thereof. Preferably, the first and second amino acid sequences in the fusion polypeptide comprise different epitopes.

Hybrids (or fusions) consisting of amino acid sequences from two, three, four, five, six, seven, eight, nine, or ten GBS antigens are preferred. In particular, hybrids consisting of amino acid sequences from two, three, four, or five GBS antigens are preferred.

Different hybrid polypeptides may be mixed together in a single formulation. Within such combinations, a GBS antigen may be present in more than one hybrid polypeptide and/or as a non-hybrid polypeptide. It is preferred, however, that an antigen is present either as a hybrid or as a non-hybrid, but not as both.

Hybrid polypeptides can be represented by the formula  $\text{NH}_2\text{-A-}\{-\text{X-L-}\}_n\text{-B-COOH}$ , wherein: X is an amino acid sequence of a GBS AI protein or a fragment thereof; L is an optional linker amino acid sequence; A is an optional N-terminal amino acid sequence; B is an optional C-terminal amino acid sequence; and  $n$  is 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14 or 15.

If a -X- moiety has a leader peptide sequence in its wild-type form, this may be included or omitted in the hybrid protein. In some embodiments, the leader peptides will be deleted except for that of the -X- moiety located at the N-terminus of the hybrid protein *i.e.* the leader peptide of  $X_1$  will be retained, but the leader peptides of  $X_2 \dots X_n$  will be omitted. This is equivalent to deleting all leader peptides and using the leader peptide of  $X_1$  as moiety -A-.

For each  $n$  instances of  $\{-\text{X-L-}\}$ , linker amino acid sequence -L- may be present or absent. For instance, when  $n=2$  the hybrid may be  $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-L}_2\text{-COOH}$ ,  $\text{NH}_2\text{-X}_1\text{-X}_2\text{-COOH}$ ,  $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-COOH}$ ,  $\text{NH}_2\text{-X}_1\text{-X}_2\text{-L}_2\text{-COOH}$ , *etc.* Linker amino acid sequence(s) -L- will typically be short (*e.g.* 20 or fewer amino acids *i.e.* 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples comprise short peptide sequences which facilitate cloning, poly-glycine linkers (*i.e.* comprising  $\text{Gly}_n$  where  $n = 2, 3, 4, 5, 6, 7, 8, 9, 10$  or more), and histidine tags (*i.e.*  $\text{His}_n$  where  $n = 3, 4, 5, 6, 7, 8, 9, 10$  or more). Other suitable linker amino acid sequences will be apparent to those skilled in the art. A useful linker is GSGGGG, with the Gly-Ser dipeptide being formed from a *Bam*HI restriction site, thus aiding cloning and manipulation, and the  $(\text{Gly})_4$  tetrapeptide being a typical poly-glycine linker.

-A- is an optional N-terminal amino acid sequence. This will typically be short (*e.g.* 40 or fewer amino acids *i.e.* 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19,



18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include leader sequences to direct protein trafficking, or short peptide sequences which facilitate cloning or purification (*e.g.* histidine tags *i.e.* His<sub>*n*</sub> where *n* = 3, 4, 5, 6, 7, 8, 9, 10 or more). Other suitable N-terminal amino acid sequences will be apparent to those skilled in the art. If X<sub>1</sub> lacks its own N-terminus methionine, -A- is preferably an oligopeptide (*e.g.* with 1, 2, 3, 4, 5, 6, 7 or 8 amino acids) which provides a N-terminus methionine.

-B- is an optional C-terminal amino acid sequence. This will typically be short (*e.g.* 40 or fewer amino acids *i.e.* 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include sequences to direct protein trafficking, short peptide sequences which facilitate cloning or purification (*e.g.* comprising histidine tags *i.e.* His<sub>*n*</sub> where *n* = 3, 4, 5, 6, 7, 8, 9, 10 or more), or sequences which enhance protein stability. Other suitable C-terminal amino acid sequences will be apparent to those skilled in the art.

Most preferably, *n* is 2 or 3.

#### Fusion Proteins: Gram positive bacteria AI sequences

The Gram positive bacteria AI proteins used in the invention may be present in the composition as individual separate polypeptides, but it is preferred that at least two (*i.e.* 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, or 18) of the antigens are expressed as a single polypeptide chain (a "hybrid" or "fusion" polypeptide). Such fusion polypeptides offer two principal advantages: first, a polypeptide that may be unstable or poorly expressed on its own can be assisted by adding a suitable fusion partner that overcomes the problem; second, commercial manufacture is simplified as only one expression and purification need be employed in order to produce two polypeptides which are both antigenically useful.

The fusion polypeptide may comprise one or more AI polypeptide sequences. Preferably, the fusion comprises an AI surface protein sequence. Accordingly, the invention includes a fusion peptide comprising a first amino acid sequence and a second amino acid sequence, wherein said first and second amino acid sequences are selected from a Gram positive bacteria AI protein or a fragment thereof. Preferably, the first and second amino acid sequences in the fusion polypeptide comprise different epitopes.

Hybrids (or fusions) consisting of amino acid sequences from two, three, four, five, six, seven, eight, nine, or ten Gram positive bacteria antigens are preferred. In particular, hybrids consisting of amino acid sequences from two, three, four, or five Gram positive bacteria antigens are preferred.

Different hybrid polypeptides may be mixed together in a single formulation. Within such combinations, a Gram positive bacteria AI sequence may be present in more than one hybrid polypeptide and/or as a non-hybrid polypeptide. It is preferred, however, that an antigen is present either as a hybrid or as a non-hybrid, but not as both.

Hybrid polypeptides can be represented by the formula NH<sub>2</sub>-A-{-X-L-}<sub>*n*</sub>-B-COOH, wherein: X is an amino acid sequence of a Gram positive bacteria AI sequence or a fragment thereof; L is an

optional linker amino acid sequence; A is an optional N-terminal amino acid sequence; B is an optional C-terminal amino acid sequence; and  $n$  is 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14 or 15.

If a -X- moiety has a leader peptide sequence in its wild-type form, this may be included or omitted in the hybrid protein. In some embodiments, the leader peptides will be deleted except for that of the -X- moiety located at the N-terminus of the hybrid protein *i.e.* the leader peptide of  $X_1$  will be retained, but the leader peptides of  $X_2 \dots X_n$  will be omitted. This is equivalent to deleting all leader peptides and using the leader peptide of  $X_1$  as moiety -A-.

For each  $n$  instances of {-X-L-}, linker amino acid sequence -L- may be present or absent. For instance, when  $n=2$  the hybrid may be  $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-L}_2\text{-COOH}$ ,  $\text{NH}_2\text{-X}_1\text{-X}_2\text{-COOH}$ ,  $\text{NH}_2\text{-X}_1\text{-L}_1\text{-X}_2\text{-COOH}$ ,  $\text{NH}_2\text{-X}_1\text{-X}_2\text{-L}_2\text{-COOH}$ , *etc.* Linker amino acid sequence(s) -L- will typically be short (*e.g.* 20 or fewer amino acids *i.e.* 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples comprise short peptide sequences which facilitate cloning, poly-glycine linkers (*i.e.* comprising  $\text{Gly}_n$  where  $n = 2, 3, 4, 5, 6, 7, 8, 9, 10$  or more), and histidine tags (*i.e.*  $\text{His}_n$  where  $n = 3, 4, 5, 6, 7, 8, 9, 10$  or more). Other suitable linker amino acid sequences will be apparent to those skilled in the art. A useful linker is GSGGGG, with the Gly-Ser dipeptide being formed from a *Bam*HI restriction site, thus aiding cloning and manipulation, and the  $(\text{Gly})_4$  tetrapeptide being a typical poly-glycine linker.

-A- is an optional N-terminal amino acid sequence. This will typically be short (*e.g.* 40 or fewer amino acids *i.e.* 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include leader sequences to direct protein trafficking, or short peptide sequences which facilitate cloning or purification (*e.g.* histidine tags *i.e.*  $\text{His}_n$  where  $n = 3, 4, 5, 6, 7, 8, 9, 10$  or more). Other suitable N-terminal amino acid sequences will be apparent to those skilled in the art. If  $X_1$  lacks its own N-terminus methionine, -A- is preferably an oligopeptide (*e.g.* with 1, 2, 3, 4, 5, 6, 7 or 8 amino acids) which provides a N-terminus methionine.

-B- is an optional C-terminal amino acid sequence. This will typically be short (*e.g.* 40 or fewer amino acids *i.e.* 39, 38, 37, 36, 35, 34, 33, 32, 31, 30, 29, 28, 27, 26, 25, 24, 23, 22, 21, 20, 19, 18, 17, 16, 15, 14, 13, 12, 11, 10, 9, 8, 7, 6, 5, 4, 3, 2, 1). Examples include sequences to direct protein trafficking, short peptide sequences which facilitate cloning or purification (*e.g.* comprising histidine tags *i.e.*  $\text{His}_n$  where  $n = 3, 4, 5, 6, 7, 8, 9, 10$  or more), or sequences which enhance protein stability. Other suitable C-terminal amino acid sequences will be apparent to those skilled in the art.

Most preferably,  $n$  is 2 or 3.

#### Antibodies: GBS AI sequences

The GBS AI proteins of the invention may also be used to prepare antibodies specific to the GBS AI proteins. The antibodies are preferably specific to the an oligomeric or hyper-oligomeric form of an AI protein. The invention also includes combinations of antibodies specific to GBS AI proteins selected to provide protection against an increased range of GBS serotypes and strain isolates. For example, a combination may comprise a first and second antibody, wherein said first



antibody is specific to a first GBS AI protein and said second antibody is specific to a second GBS AI protein. Preferably, the nucleic acid sequence encoding said first GBS AI protein is not present in a GBS genome comprising a polynucleotide sequence encoding for said second GBS AI protein. Preferably, the nucleic acid sequence encoding said first and second GBS AI proteins are present in the genomes of multiple GBS serotypes and strain isolates.

The GBS specific antibodies of the invention include one or more biological moieties that, through chemical or physical means, can bind to or associate with an epitope of a GBS polypeptide. The antibodies of the invention include antibodies which specifically bind to a GBS AI protein. The invention includes antibodies obtained from both polyclonal and monoclonal preparations, as well as the following: hybrid (chimeric) antibody molecules (see, for example, Winter *et al.* (1991) *Nature* 349: 293-299; and US Patent No. 4,816,567; F(ab')<sub>2</sub> and F(ab) fragments; F<sub>v</sub> molecules (non-covalent heterodimers, see, for example, Inbar *et al.* (1972) *Proc Natl Acad Sci USA* 69:2659-2662; and Ehrlich *et al.* (1980) *Biochem* 19:4091-4096); single-chain Fv molecules (sFv) (see, for example, Huston *et al.* (1988) *Proc Natl Acad Sci USA* 85:5897-5883); dimeric and trimeric antibody fragment constructs; minibodies (see, *e.g.*, Pack *et al.* (1992) *Biochem* 31:1579-1584; Cumber *et al.* (1992) *J Immunology* 149B: 120-126); humanized antibody molecules (see, for example, Riechmann *et al.* (1988) *Nature* 332:323-327; Verhoeyan *et al.* (1988) *Science* 239:1534-1536; and U.K. Patent Publication No. GB 2,276,169, published 21 September 1994); and, any functional fragments obtained from such molecules, wherein such fragments retain immunological binding properties of the parent antibody molecule. The invention further includes antibodies obtained through non-conventional processes, such as phage display.

Preferably, the GBS specific antibodies of the invention are monoclonal antibodies. Monoclonal antibodies of the invention include an antibody composition having a homogeneous antibody population. Monoclonal antibodies of the invention may be obtained from murine hybridomas, as well as human monoclonal antibodies obtained using human rather than murine hybridomas. See, *e.g.*, Cote, *et al. Monoclonal Antibodies and Cancer Therapy*, Alan R. Liss, 1985, p 77.

The antibodies of the invention may be used in diagnostic applications, for example, to detect the presence or absence of GBS in a biological sample. The antibodies of the invention may also be used in the prophylactic or therapeutic treatment of GBS infection.

#### Antibodies: Gram positive bacteria AI sequences

The Gram positive bacteria AI proteins of the invention may also be used to prepare antibodies specific to the Gram positive bacteria AI proteins. The antibodies are preferably specific to the an oligomeric or hyper-oligomeric form of an AI protein. The invention also includes combinations of antibodies specific to Gram positive bacteria AI proteins selected to provide protection against an increased range of Gram positive bacteria genus, species, serotypes and strain isolates.

For example, a combination may comprise a first and second antibody, wherein said first antibody is specific to a first Gram positive bacteria AI protein and said second antibody is specific to a second Gram positive bacteria AI protein. Preferably, the nucleic acid sequence encoding said first Gram positive bacteria AI protein is not present in a Gram positive bacterial genome comprising a polynucleotide sequence encoding for said second Gram positive bacteria AI protein. Preferably, the nucleic acid sequence encoding said first and second Gram positive bacteria AI proteins are present in the genomes of multiple Gram positive bacteria genus, species, serotypes or strain isolates.

As an example of an instance where the combination of antibodies provides protection against an increased range of bacteria serotypes, the first antibody may be specific to a first GAS AI protein and the second antibody may be specific to a second GAS AI protein. The first GAS AI protein may comprise a GAS AI-1 surface protein, while the second GAS AI protein may comprise a GAS AI-2 or AI-3 surface protein.

As an example of an instance where the combination of antibodies provides protection against an increased range of bacterial species, the first antibody may be specific to a GBS AI protein and the second antibody may be specific to a GAS AI protein. Alternatively, the first antibody may be specific to a GAS AI protein and the second antibody may be specific to a *S. pneumoniae* AI protein.

The Gram positive specific antibodies of the invention include one or more biological moieties that, through chemical or physical means, can bind to or associate with an epitope of a Gram positive bacteria AI polypeptide. The antibodies of the invention include antibodies which specifically bind to a Gram positive bacteria AI protein. The invention includes antibodies obtained from both polyclonal and monoclonal preparations, as well as the following: hybrid (chimeric) antibody molecules (see, for example, Winter *et al.* (1991) *Nature* 349: 293-299; and US Patent No. 4,816,567; F(ab')<sub>2</sub> and F(ab) fragments; F<sub>v</sub> molecules (non-covalent heterodimers, see, for example, Inbar *et al.* (1972) *Proc Natl Acad Sci USA* 69:2659-2662; and Ehrlich *et al.* (1980) *Biochem* 19:4091-4096); single-chain F<sub>v</sub> molecules (sFv) (see, for example, Huston *et al.* (1988) *Proc Natl Acad Sci USA* 85:5897-5883); dimeric and trimeric antibody fragment constructs; minibodies (see, *e.g.*, Pack *et al.* (1992) *Biochem* 31:1579-1584; Cumber *et al.* (1992) *J Immunology* 149B: 120-126); humanized antibody molecules (see, for example, Riechmann *et al.* (1988) *Nature* 332:323-327; Verhoeyan *et al.* (1988) *Science* 239:1534-1536; and U.K. Patent Publication No. GB 2,276,169, published 21 September 1994); and, any functional fragments obtained from such molecules, wherein such fragments retain immunological binding properties of the parent antibody molecule. The invention further includes antibodies obtained through non-conventional processes, such as phage display.

Preferably, the Gram positive specific antibodies of the invention are monoclonal antibodies. Monoclonal antibodies of the invention include an antibody composition having a homogeneous antibody population. Monoclonal antibodies of the invention may be obtained from murine hybridomas, as well as human monoclonal antibodies obtained using human rather than murine



The antibodies of the invention may be used in diagnostic applications, for example, to detect the presence or absence of Gram positive bacteria in a biological sample. The antibodies of the invention may also be used in the prophylactic or therapeutic treatment of Gram positive bacteria infection.

#### Nucleic Acids

The invention provides nucleic acids encoding the Gram positive bacteria sequences and/or the hybrid fusion polypeptides of the invention. The invention also provides nucleic acid encoding the GBS antigens and/or the hybrid fusion polypeptides of the invention. Furthermore, the invention provides nucleic acid which can hybridise to these nucleic acids, preferably under "high stringency" conditions (e.g. 65°C in a 0.1xSSC, 0.5% SDS solution).

Polypeptides of the invention can be prepared by various means (e.g. recombinant expression, purification from cell culture, chemical synthesis, *etc.*) and in various forms (e.g. native, fusions, non-glycosylated, lipidated, *etc.*). They are preferably prepared in substantially pure form (*i.e.* substantially free from other GAS or host cell proteins).

Nucleic acid according to the invention can be prepared in many ways (e.g. by chemical synthesis, from genomic or cDNA libraries, from the organism itself, *etc.*) and can take various forms (e.g. single stranded, double stranded, vectors, probes, *etc.*). They are preferably prepared in substantially pure form (*i.e.* substantially free from other GBS or host cell nucleic acids).

The term "nucleic acid" includes DNA and RNA, and also their analogues, such as those containing modified backbones (e.g. phosphorothioates, *etc.*), and also peptide nucleic acids (PNA), *etc.* The invention includes nucleic acid comprising sequences complementary to those described above (e.g. for antisense or probing purposes).

The invention also provides a process for producing a polypeptide of the invention, comprising the step of culturing a host cell transformed with nucleic acid of the invention under conditions which induce polypeptide expression.

The invention provides a process for producing a polypeptide of the invention, comprising the step of synthesising at least part of the polypeptide by chemical means.

The invention provides a process for producing nucleic acid of the invention, comprising the step of amplifying nucleic acid using a primer-based amplification method (e.g. PCR).

The invention provides a process for producing nucleic acid of the invention, comprising the step of synthesising at least part of the nucleic acid by chemical means.

#### Purification and Recombinant Expression

The Gram positive bacteria AI proteins of the invention may be isolated from the native Gram positive bacteria, or they may be recombinantly produced, for instance in a heterologous host. For example, the GAS, GBS, and *S. pneumoniae* antigens of the invention may be isolated from

*Streptococcus agalactiae*, *S. pyogenes*, *S. pneumoniae*, or they may be recombinantly produced, for instance, in a heterologous host. Preferably, the GBS antigens are prepared using a heterologous host.

The heterologous host may be prokaryotic (e.g. a bacterium) or eukaryotic. It is preferably *E. coli*, but other suitable hosts include *Bacillus subtilis*, *Vibrio cholerae*, *Salmonella typhi*, *Salmonella typhimurium*, *Neisseria lactamica*, *Neisseria cinerea*, *Mycobacteria* (e.g. *M. tuberculosis*), *S. gordonii*, *L. lactis*, yeasts, etc.

Recombinant production of polypeptides is facilitated by adding a tag protein to the Gram positive bacteria AI sequence to be expressed as a fusion protein comprising the tag protein and the Gram positive bacteria antigen. For example, recombinant production of polypeptides is facilitated by adding a tag protein to the GBS antigen to be expressed as a fusion protein comprising the tag protein and the GBS antigen. Such tag proteins can facilitate purification, detection and stability of the expressed protein. Tag proteins suitable for use in the invention include a polyarginine tag (Arg-tag), polyhistidine tag (His-tag), FLAG-tag, Strep-tag, c-myc-tag, S-tag, calmodulin-binding peptide, cellulose-binding domain, SBP-tag, chitin-binding domain, glutathione S-transferase-tag (GST), maltose-binding protein, transcription termination anti-terminiation factor (NusA), *E. coli* thioredoxin (TrxA) and protein disulfide isomerase I (DsbA). Preferred tag proteins include His-tag and GST. A full discussion on the use of tag proteins can be found at Terpe et al., "Overview of tag protein fusions: from molecular and biochemical fundamentals to commercial systems", Appl Microbiol Biotechnol (2003) 60:523 – 533.

After purification, the tag proteins may optionally be removed from the expressed fusion protein, i.e., by specifically tailored enzymatic treatments known in the art. Commonly used proteases include enterokinase, tobacco etch virus (TEV), thrombin, and factor X<sub>a</sub>.

#### GBS polysaccharides

The compositions of the invention may be further improved by including GBS polysaccharides. Preferably, the GBS antigen and the saccharide each contribute to the immunological response in a recipient. The combination is particularly advantageous where the saccharide and polypeptide provide protection from different GBS serotypes.

The combined antigens may be present as a simple combination where separate saccharide and polypeptide antigens are administered together, or they may be present as a conjugated combination, where the saccharide and polypeptide antigens are covalently linked to each other.

Thus the invention provides an immunogenic composition comprising (i) one or more GBS AI proteins and (ii) one or more GBS saccharide antigens. The polypeptide and the polysaccharide may advantageously be covalently linked to each other to form a conjugate.

Between them, the combined polypeptide and saccharide antigens preferably cover (or provide protection from) two or more GBS serotypes (e.g. 2, 3, 4, 5, 6, 7, 8 or more serotypes). The serotypes of the polypeptide and saccharide antigens may or may not overlap. For example, the polypeptide might protect against serogroup II or V, while the saccharide protects against either serogroups Ia, Ib, or III. Preferred combinations protect against the following groups of serotypes:



(1) serotypes Ia and Ib, (2) serotypes Ia and II, (3) serotypes Ia and III, (4) serotypes Ia and IV, (5) serotypes Ia and V, (6) serotypes Ia and VI, (7) serotypes Ia and VII, (8) serotypes Ia and VIII, (9) serotypes Ib and II, (10) serotypes Ib and III, (11) serotypes Ib and IV, (12) serotypes Ib and V, (13) serotypes Ib and VI, (14) serotypes Ib and VII, (15) serotypes Ib and VIII, 16) serotypes II and III, 5 (17) serotypes II and IV, (18) serotypes II and V, (19) serotypes II and VI, (20) serotypes II and VII, (21) serotypes II and VIII, (22) serotypes III and IV, (23) serotypes III and V, (24) serotypes III and VI, (25) serotypes III and VII, (26) serotypes III and VIII, (27) serotypes IV and V, (28) serotypes IV and VI, (29) serotypes IV and VII, (30) serotypes IV and VIII, (31) serotypes V and VI, (32) serotypes V and VII, (33) serotypes V and VIII, (34) serotypes VI and VII, (35) serotypes VI and 10 VIII, and (36) serotypes VII and VIII.

Still more preferably, the combinations protect against the following groups of serotypes: (1) serotypes Ia and II, (2) serotypes Ia and V, (3) serotypes Ib and II, (4) serotypes Ib and V, (5) serotypes III and II, and (6) serotypes III and V. Most preferably, the combinations protect against serotypes III and V.

15 Protection against serotypes II and V is preferably provided by polypeptide antigens. Protection against serotypes Ia, Ib and/or III may be polypeptide or saccharide antigens.

#### Immunogenic compositions and medicaments

Compositions of the invention are preferably immunogenic compositions, and are more preferably vaccine compositions. The pH of the composition is preferably between 6 and 8, 20 preferably about 7. The pH may be maintained by the use of a buffer. The composition may be sterile and/or pyrogen-free. The composition may be isotonic with respect to humans.

Vaccines according to the invention may either be prophylactic (*i.e.* to prevent infection) or therapeutic (*i.e.* to treat infection), but will typically be prophylactic. Accordingly, the invention includes a method for the therapeutic or prophylactic treatment of a Gram positive bacteria infection 25 in an animal susceptible to such gram positive bacterial infection comprising administering to said animal a therapeutic or prophylactic amount of the immunogenic composition of the invention. For example, the invention includes a method for the therapeutic or prophylactic treatment of a *Streptococcus agalactiae*, *S. pyogenes*, or *S. pneumoniae* infection in an animal susceptible to streptococcal infection comprising administering to said animal a therapeutic or prophylactic amount 30 of the immunogenic compositions of the invention.

The invention also provides a composition of the invention for use of the compositions described herein as a medicament. The medicament is preferably able to raise an immune response in a mammal (*i.e.* it is an immunogenic composition) and is more preferably a vaccine.

35 The invention also provides the use of the compositions of the invention in the manufacture of a medicament for raising an immune response in a mammal. The medicament is preferably a vaccine.

The invention also provides kits comprising one or more containers of compositions of the invention. Compositions can be in liquid form or can be lyophilized, as can individual antigens. Suitable containers for the compositions include, for example, bottles, vials, syringes, and test tubes.

Containers can be formed from a variety of materials, including glass or plastic. A container may have a sterile access port (for example, the container may be an intravenous solution bag or a vial having a stopper pierceable by a hypodermic injection needle). The composition may comprise a first component comprising one or more Gram positive bacteria AI proteins. Preferably, the AI proteins are surface AI proteins. Preferably, the AI surface proteins are in an oligomeric or hyperoligomeric form. For example, the first component comprises a combination of GBS antigens or GAS antigens, or *S. pneumoniae* antigens. Preferably said combination includes GBS 80. Preferably GBS 80 is present in an oligomeric or hyperoligomeric form.

The kit can further comprise a second container comprising a pharmaceutically-acceptable buffer, such as phosphate-buffered saline, Ringer's solution, or dextrose solution. It can also contain other materials useful to the end-user, including other buffers, diluents, filters, needles, and syringes. The kit can also comprise a second or third container with another active agent, for example an antibiotic.

The kit can also comprise a package insert containing written instructions for methods of inducing immunity against *S. agalactiae* and/or *S. pyogenes* and/or *S. pneumoniae* or for treating *S. agalactiae* and/or *S. pyogenes* and/or *S. pneumoniae* infections. The package insert can be an unapproved draft package insert or can be a package insert approved by the Food and Drug Administration (FDA) or other regulatory body.

The invention also provides a delivery device pre-filled with the immunogenic compositions of the invention.

The invention also provides a method for raising an immune response in a mammal comprising the step of administering an effective amount of a composition of the invention. The immune response is preferably protective and preferably involves antibodies and/or cell-mediated immunity. This immune response will preferably induce long lasting (e.g., neutralising) antibodies and a cell mediated immunity that can quickly respond upon exposure to one or more GBS and/or GAS and/or *S. pneumoniae* antigens. The method may raise a booster response.

The invention provides a method of neutralizing GBS, GAS, or *S. pneumoniae* infection in a mammal comprising the step of administering to the mammal an effective amount of the immunogenic compositions of the invention, a vaccine of the invention, or antibodies which recognize an immunogenic composition of the invention.

The mammal is preferably a human. Where the vaccine is for prophylactic use, the human is preferably a female (either of child bearing age or a teenager). Alternatively, the human may be elderly (e.g., over the age of 50, 55, 60, 65, 70 or 75) and may have an underlying disease such as diabetes or cancer. Where the vaccine is for therapeutic use, the human is preferably a pregnant female or an elderly adult.

These uses and methods are preferably for the prevention and/or treatment of a disease caused by *Streptococcus agalactiae*, or *S. pyogenes*, or *S. pneumoniae*. The compositions may also be



effective against other streptococcal bacteria. The compositions may also be effective against other Gram positive bacteria.

One way of checking efficacy of therapeutic treatment involves monitoring Gram positive bacterial infection after administration of the composition of the invention. One way of checking efficacy of prophylactic treatment involves monitoring immune responses against the Gram positive bacterial antigens in the compositions of the invention after administration of the composition.

One way of checking efficacy of therapeutic treatment involves monitoring GBS infection after administration of the composition of the invention. One way of checking efficacy of prophylactic treatment involves monitoring immune responses against the GBS antigens in the compositions of the invention after administration of the composition.

A way of assessing the immunogenicity of the component proteins of the immunogenic compositions of the present invention is to express the proteins recombinantly and to screen patient sera or mucosal secretions by immunoblot. A positive reaction between the protein and the patient serum indicates that the patient has previously mounted an immune response to the protein in question- that is, the protein is an immunogen. This method may also be used to identify immunodominant proteins and/or epitopes.

Another way of checking efficacy of therapeutic treatment involves monitoring GBS or GAS or *S pneumoniae* infection after administration of the compositions of the invention. One way of checking efficacy of prophylactic treatment involves monitoring immune responses both systemically (such as monitoring the level of IgG1 and IgG2a production) and mucosally (such as monitoring the level of IgA production) against the GBS and/or GAS and/or *S pneumoniae* antigens in the compositions of the invention after administration of the composition. Typically, GBS and/or GAS and/or *S pneumoniae* serum specific antibody responses are determined post-immunization but pre-challenge whereas mucosal GBS and/or GAS and/or *S pneumoniae* specific antibody body responses are determined post-immunization and post-challenge.

The vaccine compositions of the present invention can be evaluated in *in vitro* and *in vivo* animal models prior to host, e.g., human, administration.

The efficacy of immunogenic compositions of the invention can also be determined *in vivo* by challenging animal models of GBS and/or GAS and/or *S pneumoniae* infection, e.g., guinea pigs or mice, with the immunogenic compositions. The immunogenic compositions may or may not be derived from the same serotypes as the challenge serotypes. Preferably the immunogenic compositions are derivable from the same serotypes as the challenge serotypes. More preferably, the immunogenic composition and/or the challenge serotypes are derivable from the group of GBS and/or GAS and/or *S pneumoniae* serotypes.

*In vivo* efficacy models include but are not limited to: (i) A murine infection model using human GBS and/or GAS and/or *S pneumoniae* serotypes; (ii) a murine disease model which is a murine model using a mouse-adapted GBS and/or GAS and/or *S pneumoniae* strain, such as those

strains outlined above which is particularly virulent in mice and (iii) a primate model using human GBS or GAS or S pneumoniae isolates.

The immune response may be one or both of a TH1 immune response and a TH2 response.

The immune response may be an improved or an enhanced or an altered immune response.

5 The immune response may be one or both of a systemic and a mucosal immune response.

Preferably the immune response is an enhanced system and/or mucosal response.

An enhanced systemic and/or mucosal immunity is reflected in an enhanced TH1 and/or TH2 immune response. Preferably, the enhanced immune response includes an increase in the production of IgG1 and/or IgG2a and/or IgA

10 Preferably the mucosal immune response is a TH2 immune response. Preferably, the mucosal immune response includes an increase in the production of IgA.

Activated TH2 cells enhance antibody production and are therefore of value in responding to extracellular infections. Activated TH2 cells may secrete one or more of IL-4, IL-5, IL-6, and IL-10.

15 A TH2 immune response may result in the production of IgG1, IgE, IgA and memory B cells for future protection.

A TH2 immune response may include one or more of an increase in one or more of the cytokines associated with a TH2 immune response (such as IL-4, IL-5, IL-6 and IL-10), or an increase in the production of IgG1, IgE, IgA and memory B cells. Preferably, the enhanced TH2 immune response will include an increase in IgG1 production.

20 A TH1 immune response may include one or more of an increase in CTLs, an increase in one or more of the cytokines associated with a TH1 immune response (such as IL-2, IFN $\gamma$ , and TNF $\beta$ ), an increase in activated macrophages, an increase in NK activity, or an increase in the production of IgG2a. Preferably, the enhanced TH1 immune response will include an increase in IgG2a production.

25 Immunogenic compositions of the invention, in particular, immunogenic composition comprising one or more GAS antigens of the present invention may be used either alone or in combination with other GAS antigens optionally with an immunoregulatory agent capable of eliciting a Th1 and/or Th2 response.

30 Compositions of the invention will generally be administered directly to a patient. Certain routes may be favored for certain compositions, as resulting in the generation of a more effective immune response, preferably a CMI response, or as being less likely to induce side effects, or as being easier for administration. Direct delivery may be accomplished by parenteral injection (*e.g.* subcutaneously, intraperitoneally, intradermally, intravenously, intramuscularly, or to the interstitial space of a tissue), or by rectal, oral (*e.g.* tablet, spray), vaginal, topical, transdermal (*e.g.* see WO 99/27961) or transcutaneous (*e.g.* see WO 02/074244 and WO 02/064162), intranasal (*e.g.* see 35 WO03/028760), ocular, aural, pulmonary or other mucosal administration.

The invention may be used to elicit systemic and/or mucosal immunity.



In one particularly preferred embodiment, the immunogenic composition comprises one or more GBS or GAS or S pneumoniae antigen(s) which elicits a neutralising antibody response and one or more GBS or GAS or S pneumoniae antigen(s) which elicit a cell mediated immune response. In this way, the neutralising antibody response prevents or inhibits an initial GBS or GAS or S

pneumoniae infection while the cell-mediated immune response capable of eliciting an enhanced Th1 cellular response prevents further spreading of the GBS or GAS or S pneumoniae infection.

Preferably, the immunogenic composition comprises one or more GBS or GAS or S pneumoniae surface antigens and one or more GBS or GAS or S pneumoniae cytoplasmic antigens. Preferably the immunogenic composition comprises one or more GBS or GAS or S pneumoniae surface antigens or the like and one or other antigens, such as a cytoplasmic antigen capable of eliciting a Th1 cellular response.

Dosage treatment can be a single dose schedule or a multiple dose schedule. Multiple doses may be used in a primary immunisation schedule and/or in a booster immunisation schedule. In a multiple dose schedule the various doses may be given by the same or different routes *e.g.* a

parenteral prime and mucosal boost, a mucosal prime and parenteral boost, *etc.*

The compositions of the invention may be prepared in various forms. For example, the compositions may be prepared as injectables, either as liquid solutions or suspensions. Solid forms suitable for solution in, or suspension in, liquid vehicles prior to injection can also be prepared (*e.g.* a lyophilised composition). The composition may be prepared for topical administration *e.g.* as an ointment, cream or powder. The composition may be prepared for oral administration *e.g.* as a tablet or capsule, as a spray, or as a syrup (optionally flavoured). The composition may be prepared for pulmonary administration *e.g.* as an inhaler, using a fine powder or a spray. The composition may be prepared as a suppository or pessary. The composition may be prepared for nasal, aural or ocular administration *e.g.* as drops. The composition may be in kit form, designed such that a combined composition is reconstituted just prior to administration to a patient. Such kits may comprise one or more antigens in liquid form and one or more lyophilised antigens.

Immunogenic compositions used as vaccines comprise an immunologically effective amount of antigen(s), as well as any other components, such as antibiotics, as needed. By 'immunologically effective amount', it is meant that the administration of that amount to an individual, either in a single dose or as part of a series, is effective for treatment or prevention, or increases a measurable immune response or prevents or reduces a clinical symptom. This amount varies depending upon the health and physical condition of the individual to be treated, age, the taxonomic group of individual to be treated (*e.g.* non-human primate, primate, *etc.*), the capacity of the individual's immune system to synthesise antibodies, the degree of protection desired, the formulation of the vaccine, the treating doctor's assessment of the medical situation, and other relevant factors. It is expected that the amount will fall in a relatively broad range that can be determined through routine trials.

Further Components of the Composition

The composition of the invention will typically, in addition to the components mentioned above, comprise one or more 'pharmaceutically acceptable carriers', which include any carrier that does not itself induce the production of antibodies harmful to the individual receiving the composition. Suitable carriers are typically large, slowly metabolised macromolecules such as proteins, polysaccharides, polylactic acids, polyglycolic acids, polymeric amino acids, amino acid copolymers, and lipid aggregates (such as oil droplets or liposomes). Such carriers are well known to those of ordinary skill in the art. The vaccines may also contain diluents, such as water, saline, glycerol, *etc.* Additionally, auxiliary substances, such as wetting or emulsifying agents, pH buffering substances, and the like, may be present. A thorough discussion of pharmaceutically acceptable excipients is available in Gennaro (2000) *Remington: The Science and Practice of Pharmacy*. 20th ed., ISBN: 0683306472.

Adjuvants

Vaccines of the invention may be administered in conjunction with other immunoregulatory agents. In particular, compositions will usually include an adjuvant. Adjuvants for use with the invention include, but are not limited to, one or more of the following set forth below:

*A. Mineral Containing Compositions*

Mineral containing compositions suitable for use as adjuvants in the invention include mineral salts, such as aluminum salts and calcium salts. The invention includes mineral salts such as hydroxides (*e.g.* oxyhydroxides), phosphates (*e.g.* hydroxyphosphates, orthophosphates), sulfates, *etc.* (*e.g.* see chapters 8 & 9 of *Vaccine Design...* (1995) eds. Powell & Newman. ISBN: 030644867X. Plenum.), or mixtures of different mineral compounds (*e.g.* a mixture of a phosphate and a hydroxide adjuvant, optionally with an excess of the phosphate), with the compounds taking any suitable form (*e.g.* gel, crystalline, amorphous, *etc.*), and with adsorption to the salt(s) being preferred. The mineral containing compositions may also be formulated as a particle of metal salt (WO 00/23105).

Aluminum salts may be included in vaccines of the invention such that the dose of  $Al^{3+}$  is between 0.2 and 1.0 mg per dose.

*B. Oil-Emulsions*

Oil-emulsion compositions suitable for use as adjuvants in the invention include squalene-water emulsions, such as MF59 (5% Squalene, 0.5% Tween 80, and 0.5% Span 85, formulated into submicron particles using a microfluidizer). See WO90/14837. See also, Podda, "The adjuvanted influenza vaccines with novel adjuvants: experience with the MF59-adjuvanted vaccine", *Vaccine* (2001) 19: 2673-2680; Frey et al., "Comparison of the safety, tolerability, and immunogenicity of a MF59-adjuvanted influenza vaccine and a non-adjuvanted influenza vaccine in non-elderly adults", *Vaccine* (2003) 21:4234-4237. MF59 is used as the adjuvant in the FLUAD™ influenza virus trivalent subunit vaccine.



Particularly preferred adjuvants for use in the compositions are submicron oil-in-water emulsions. Preferred submicron oil-in-water emulsions for use herein are squalene/water emulsions optionally containing varying amounts of MTP-PE, such as a submicron oil-in-water emulsion containing 4-5% w/v squalene, 0.25-1.0% w/v Tween 80™ (polyoxyethylsorbitan monooleate), and/or 0.25-1.0% Span 85™ (sorbitan trioleate), and, optionally, N-acetylmuramyl-L-alanyl-D-isoglutaminyl-L-alanine-2-(1'-2'-dipalmitoyl-*sn*-glycero-3-hydroxyphosphoryloxy)-ethylamine (MTP-PE), for example, the submicron oil-in-water emulsion known as "MF59" (International Publication No. WO 90/14837; US Patent Nos. 6,299,884 and 6,451,325, incorporated herein by reference in their entireties; and Ott et al., "MF59 -- Design and Evaluation of a Safe and Potent Adjuvant for Human Vaccines" in *Vaccine Design: The Subunit and Adjuvant Approach* (Powell, M.F. and Newman, M.J. eds.) Plenum Press, New York, 1995, pp. 277-296). MF59 contains 4-5% w/v Squalene (e.g. 4.3%), 0.25-0.5% w/v Tween 80™, and 0.5% w/v Span 85™ and optionally contains various amounts of MTP-PE, formulated into submicron particles using a microfluidizer such as Model 110Y microfluidizer (Microfluidics, Newton, MA). For example, MTP-PE may be present in an amount of about 0-500 µg/dose, more preferably 0-250 µg/dose and most preferably, 0-100 µg/dose. As used herein, the term "MF59-0" refers to the above submicron oil-in-water emulsion lacking MTP-PE, while the term MF59-MTP denotes a formulation that contains MTP-PE. For instance, "MF59-100" contains 100 µg MTP-PE per dose, and so on. MF69, another submicron oil-in-water emulsion for use herein, contains 4.3% w/v squalene, 0.25% w/v Tween 80™, and 0.75% w/v Span 85™ and optionally MTP-PE. Yet another submicron oil-in-water emulsion is MF75, also known as SAF, containing 10% squalene, 0.4% Tween 80™, 5% pluronic-blocked polymer L121, and thr-MDP, also microfluidized into a submicron emulsion. MF75-MTP denotes an MF75 formulation that includes MTP, such as from 100-400 µg MTP-PE per dose.

Submicron oil-in-water emulsions, methods of making the same and immunostimulating agents, such as muramyl peptides, for use in the compositions, are described in detail in International Publication No. WO 90/14837 and US Patent Nos. 6,299,884 and 6,451,325, incorporated herein by reference in their entireties.

Complete Freund's adjuvant (CFA) and incomplete Freund's adjuvant (IFA) may also be used as adjuvants in the invention.

#### C. Saponin Formulations

Saponin formulations, may also be used as adjuvants in the invention. Saponins are a heterologous group of sterol glycosides and triterpenoid glycosides that are found in the bark, leaves, stems, roots and even flowers of a wide range of plant species. Saponin from the bark of the *Quillaja saponaria* Molina tree have been widely studied as adjuvants. Saponin can also be commercially obtained from *Smilax ornata* (sarsapilla), *Gypsophilla paniculata* (brides veil), and *Saponaria officianalis* (soap root). Saponin adjuvant formulations include purified formulations, such as QS21, as well as lipid formulations, such as ISCOMs.

Saponin compositions have been purified using High Performance Thin Layer Chromatography (HP-LC) and Reversed Phase High Performance Liquid Chromatography (RP-HPLC). Specific purified fractions using these techniques have been identified, including QS7, QS17, QS18, QS21, QH-A, QH-B and QH-C. Preferably, the saponin is QS21. A method of production of QS21 is disclosed in US Patent No. 5,057,540. Saponin formulations may also comprise a sterol, such as cholesterol (see WO96/33739).

Combinations of saponins and cholesterol can be used to form unique particles called Immunostimulating Complexs (ISCOMs). ISCOMs typically also include a phospholipid such as phosphatidylethanolamine or phosphatidylcholine. Any known saponin can be used in ISCOMs. Preferably, the ISCOM includes one or more of Quil A, QHA and QHC. ISCOMs are further described in EP0109942, WO 96/11711 and WO 96/33739. Optionally, the ISCOMS may be devoid of additional detergent. See WO 00/07621.

A review of the development of saponin based adjuvants can be found at Barr, et al., "ISCOMs and other saponin based adjuvants", *Advanced Drug Delivery Reviews* (1998) 32:247-271. See also Sjolander, et al., "Uptake and adjuvant activity of orally delivered saponin and ISCOM vaccines", *Advanced Drug Delivery Reviews* (1998) 32:321-338.

#### D. *Virosomes and Virus Like Particles (VLPs)*

Virosomes and Virus Like Particles (VLPs) can also be used as adjuvants in the invention. These structures generally contain one or more proteins from a virus optionally combined or formulated with a phospholipid. They are generally non-pathogenic, non-replicating and generally do not contain any of the native viral genome. The viral proteins may be recombinantly produced or isolated from whole viruses. These viral proteins suitable for use in virosomes or VLPs include proteins derived from influenza virus (such as HA or NA), Hepatitis B virus (such as core or capsid proteins), Hepatitis E virus, measles virus, Sindbis virus, Rotavirus, Foot-and-Mouth Disease virus, Retrovirus, Norwalk virus, human Papilloma virus, HIV, RNA-phages, Q $\beta$ -phage (such as coat proteins), GA-phage, fr-phage, AP205 phage, and Ty (such as retrotransposon Ty protein p1). VLPs are discussed further in WO 03/024480, WO 03/024481, and Niikura et al., "Chimeric Recombinant Hepatitis E Virus-Like Particles as an Oral Vaccine Vehicle Presenting Foreign Epitopes", *Virology* (2002) 293:273-280; Lenz et al., "Papillomavirus-Like Particles Induce Acute Activation of Dendritic Cells", *Journal of Immunology* (2001) 5246-5355; Pinto, et al., "Cellular Immune Responses to Human Papillomavirus (HPV)-16 L1 Healthy Volunteers Immunized with Recombinant HPV-16 L1 Virus-Like Particles", *Journal of Infectious Diseases* (2003) 188:327-338; and Gerber et al., "Human Papillomavirus Virus-Like Particles Are Efficient Oral Immunogens when Coadministered with Escherichia coli Heat-Labile Enterotoxin Mutant R192G or CpG", *Journal of Virology* (2001) 75(10):4752-4760. Virosomes are discussed further in, for example, Gluck et al., "New Technology Platforms in the Development of Vaccines for the Future", *Vaccine* (2002) 20:B10-B16. Immunopotentiating reconstituted influenza virosomes (IRIV) are used as the subunit antigen



E. *Bacterial or Microbial Derivatives*

Adjuvants suitable for use in the invention include bacterial or microbial derivatives such as:

5 (1) *Non-toxic derivatives of enterobacterial lipopolysaccharide (LPS)*

Such derivatives include Monophosphoryl lipid A (MPL) and 3-O-deacylated MPL (3dMPL). 3dMPL is a mixture of 3 De-O-acylated monophosphoryl lipid A with 4, 5 or 6 acylated chains. A preferred "small particle" form of 3 De-O-acylated monophosphoryl lipid A is disclosed in EP 0 689 454. Such "small particles" of 3dMPL are small enough to be sterile filtered through a 0.22 micron  
10 membrane (see EP 0 689 454). Other non-toxic LPS derivatives include monophosphoryl lipid A mimics, such as aminoalkyl glucosaminide phosphate derivatives *e.g.* RC-529. See Johnson *et al.* (1999) *Bioorg Med Chem Lett* 9:2273-2278.

(2) *Lipid A Derivatives*

Lipid A derivatives include derivatives of lipid A from *Escherichia coli* such as OM-174. OM-174 is described for example in Meraldi *et al.*, "OM-174, a New Adjuvant with a Potential for Human Use, Induces a Protective Response with Administered with the Synthetic C-Terminal  
15 Fragment 242-310 from the circumsporozoite protein of Plasmodium berghei", *Vaccine* (2003) 21:2485-2491; and Pajak, *et al.*, "The Adjuvant OM-174 induces both the migration and maturation of murine dendritic cells in vivo", *Vaccine* (2003) 21:836-842.

20 (3) *Immunostimulatory oligonucleotides*

Immunostimulatory oligonucleotides suitable for use as adjuvants in the invention include nucleotide sequences containing a CpG motif (a sequence containing an unmethylated cytosine followed by guanosine and linked by a phosphate bond). Bacterial double stranded RNA or oligonucleotides containing palindromic or poly(dG) sequences have also been shown to be  
25 immunostimulatory.

The CpG's can include nucleotide modifications/analogues such as phosphorothioate modifications and can be double-stranded or single-stranded. Optionally, the guanosine may be replaced with an analogue such as 2'-deoxy-7-deazaguanosine. See Kandimalla, *et al.*, "Divergent synthetic nucleotide motif recognition pattern: design and development of potent immunomodulatory  
30 oligodeoxyribonucleotide agents with distinct cytokine induction profiles", *Nucleic Acids Research* (2003) 31(9): 2393-2400; WO02/26757 and WO99/62923 for examples of possible analogue substitutions. The adjuvant effect of CpG oligonucleotides is further discussed in Krieg, "CpG motifs: the active ingredient in bacterial extracts?", *Nature Medicine* (2003) 9(7): 831-835; McCluskie, *et al.*, "Parenteral and mucosal prime-boost immunization strategies in mice with hepatitis B surface antigen and CpG DNA", *FEMS Immunology and Medical Microbiology* (2002) 32:179-185; WO98/40100; US Patent No. 6,207,646; US Patent No. 6,239,116 and US Patent No. 6,429,199.

The CpG sequence may be directed to TLR9, such as the motif GTCGTT or TTCGTT. See Kandimalla, *et al.*, "Toll-like receptor 9: modulation of recognition and cytokine induction by novel

synthetic CpG DNAs” Biochemical Society Transactions (2003) 31 (part 3): 654-658. The CpG sequence may be specific for inducing a Th1 immune response, such as a CpG-A ODN, or it may be more specific for inducing a B cell response, such a CpG-B ODN. CpG-A and CpG-B ODNs are discussed in Blackwell, et al., “CpG-A-Induced Monocyte IFN-gamma-Inducible Protein-10 Production is Regulated by Plasmacytoid Dendritic Cell Derived IFN-alpha”, J. Immunol. (2003) 170(8):4061-4068; Krieg, “From A to Z on CpG”, TRENDS in Immunology (2002) 23(2): 64-65 and WO01/95935. Preferably, the CpG is a CpG-A ODN.

Preferably, the CpG oligonucleotide is constructed so that the 5' end is accessible for receptor recognition. Optionally, two CpG oligonucleotide sequences may be attached at their 3' ends to form “immunomers”. See, for example, Kandimalla, et al., “Secondary structures in CpG oligonucleotides affect immunostimulatory activity”, BBRC (2003) 306:948-953; Kandimalla, et al., “Toll-like receptor 9: modulation of recognition and cytokine induction by novel synthetic CpG DNAs”, Biochemical Society Transactions (2003) 31(part 3):664-658; Bhagat et al., “CpG penta- and hexadeoxyribonucleotides as potent immunomodulatory agents” BBRC (2003) 300:853-861 and WO 03/035836.

(4) *ADP-ribosylating toxins and detoxified derivatives thereof.*

Bacterial ADP-ribosylating toxins and detoxified derivatives thereof may be used as adjuvants in the invention. Preferably, the protein is derived from *E. coli* (i.e., *E. coli* heat labile enterotoxin “LT), cholera (“CT”), or pertussis (“PT”). The use of detoxified ADP-ribosylating toxins as mucosal adjuvants is described in WO95/17211 and as parenteral adjuvants in WO98/42375. Preferably, the adjuvant is a detoxified LT mutant such as LT-K63, LT-R72, and LTR192G. The use of ADP-ribosylating toxins and detoxified derivatives thereof, particularly LT-K63 and LT-R72, as adjuvants can be found in the following references, each of which is specifically incorporated by reference herein in their entirety: Beignon, et al., “The LTR72 Mutant of Heat-Labile Enterotoxin of *Escherichia coli* Enhances the Ability of Peptide Antigens to Elicit CD4+ T Cells and Secrete Gamma Interferon after Coapplication onto Bare Skin”, Infection and Immunity (2002) 70(6):3012-3019; Pizza, et al., “Mucosal vaccines: non toxic derivatives of LT and CT as mucosal adjuvants”, Vaccine (2001) 19:2534-2541; Pizza, et al., “LTK63 and LTR72, two mucosal adjuvants ready for clinical trials” Int. J. Med. Microbiol (2000) 290(4-5):455-461; Scharton-Kersten et al., “Transcutaneous Immunization with Bacterial ADP-Ribosylating Exotoxins, Subunits and Unrelated Adjuvants”, Infection and Immunity (2000) 68(9):5306-5313; Ryan et al., “Mutants of *Escherichia coli* Heat-Labile Toxin Act as Effective Mucosal Adjuvants for Nasal Delivery of an Acellular Pertussis Vaccine: Differential Effects of the Nontoxic AB Complex and Enzyme Activity on Th1 and Th2 Cells” Infection and Immunity (1999) 67(12):6270-6280; Partidos et al., “Heat-labile enterotoxin of *Escherichia coli* and its site-directed mutant LTK63 enhance the proliferative and cytotoxic T-cell responses to intranasally co-immunized synthetic peptides”, Immunol. Lett. (1999) 67(3):209-216; Peppoloni et al., “Mutants of the *Escherichia coli* heat-labile enterotoxin as safe and strong adjuvants for intranasal delivery of vaccines”, Vaccines (2003) 2(2):285-293; and Pine et al., (2002) “Intranasal



immunization with influenza vaccine and a detoxified mutant of heat labile enterotoxin from Escherichia coli (LTK63)" J. Control Release (2002) 85(1-3):263-270. Numerical reference for amino acid substitutions is preferably based on the alignments of the A and B subunits of ADP-ribosylating toxins set forth in Domenighini et al., Mol. Microbiol (1995) 15(6):1165-1167, specifically

5 incorporated herein by reference in its entirety.

*F. Bioadhesives and Mucoadhesives*

Bioadhesives and mucoadhesives may also be used as adjuvants in the invention. Suitable bioadhesives include esterified hyaluronic acid microspheres (Singh et al. (2001) J. Cont. Rele. 70:267-276) or mucoadhesives such as cross-linked derivatives of poly(acrylic acid), polyvinyl

10 alcohol, polyvinyl pyrrolidone, polysaccharides and carboxymethylcellulose. Chitosan and derivatives thereof may also be used as adjuvants in the invention. E.g. WO99/27960.

*G. Microparticles*

Microparticles may also be used as adjuvants in the invention. Microparticles (i.e. a particle of ~100nm to ~150µm in diameter, more preferably ~200nm to ~30µm in diameter, and most

15 preferably ~500nm to ~10µm in diameter) formed from materials that are biodegradable and non-toxic (e.g. a poly(α-hydroxy acid), a polyhydroxybutyric acid, a polyorthoester, a polyanhydride, a polycaprolactone, etc.), with poly(lactide-co-glycolide) are preferred, optionally treated to have a negatively-charged surface (e.g. with SDS) or a positively-charged surface (e.g. with a cationic detergent, such as CTAB).

*H. Liposomes*

Examples of liposome formulations suitable for use as adjuvants are described in US Patent No. 6,090,406, US Patent No. 5,916,588, and EP 0 626 169.

*I. Polyoxyethylene ether and Polyoxyethylene Ester Formulations*

Adjuvants suitable for use in the invention include polyoxyethylene ethers and

25 polyoxyethylene esters. WO99/52549. Such formulations further include polyoxyethylene sorbitan ester surfactants in combination with an octoxynol (WO01/21207) as well as polyoxyethylene alkyl ethers or ester surfactants in combination with at least one additional non-ionic surfactant such as an octoxynol (WO 01/21152).

Preferred polyoxyethylene ethers are selected from the following group: polyoxyethylene-9-lauryl ether (laureth 9), polyoxyethylene-9-stearyl ether, polyoxyethylene-8-stearyl ether, polyoxyethylene-4-lauryl ether, polyoxyethylene-35-lauryl ether, and polyoxyethylene-23-lauryl ether.

*J. Polyphosphazene (PCPP)*

PCPP formulations are described, for example, in Andrianov et al., "Preparation of hydrogel

35 microspheres by coacervation of aqueous polyphosphazene solutions", Biomaterials (1998) 19(1-3):109-115 and Payne et al., "Protein Release from Polyphosphazene Matrices", Adv. Drug. Delivery Review (1998) 31(3):185-196.

*K. Muramyl peptides*

Examples of muramyl peptides suitable for use as adjuvants in the invention include N-acetyl-muramyl-L-threonyl-D-isoglutamine (thr-MDP), N-acetyl-normuramyl-l-alanyl-d-isoglutamine (nor-MDP), and N-acetylmuramyl-l-alanyl-d-isoglutaminyl-l-alanine-2-(1'-2'-dipalmitoyl-sn-glycero-3-hydroxyphosphoryloxy)-ethylamine MTP-PE).

*L. Imidazoquinolone Compounds.*

Examples of imidazoquinolone compounds suitable for use adjuvants in the invention include Imiquamod and its homologues, described further in Stanley, "Imiquimod and the imidazoquinolones: mechanism of action and therapeutic potential" Clin Exp Dermatol (2002) 27(7):571-577 and Jones, "Resiquimod 3M", Curr Opin Investig Drugs (2003) 4(2):214-218.

The invention may also comprise combinations of aspects of one or more of the adjuvants identified above. For example, the following adjuvant compositions may be used in the invention:

- (1) a saponin and an oil-in-water emulsion (WO 99/11241);
- (2) a saponin (e.g., QS21) + a non-toxic LPS derivative (e.g. 3dMPL) (see WO 94/00153);
- (3) a saponin (e.g., QS21) + a non-toxic LPS derivative (e.g. 3dMPL) + a cholesterol;
- (4) a saponin (e.g. QS21) + 3dMPL + IL-12 (optionally + a sterol) (WO 98/57659);
- (5) combinations of 3dMPL with, for example, QS21 and/or oil-in-water emulsions (See European patent applications 0835318, 0735898 and 0761231);
- (6) SAF, containing 10% Squalane, 0.4% Tween 80, 5% pluronic-block polymer L121, and thr-MDP, either microfluidized into a submicron emulsion or vortexed to generate a larger particle size emulsion.
- (7) Ribi<sup>TM</sup> adjuvant system (RAS), (Ribi Immunochem) containing 2% Squalene, 0.2% Tween 80, and one or more bacterial cell wall components from the group consisting of monophosphorylipid A (MPL), trehalose dimycolate (TDM), and cell wall skeleton (CWS), preferably MPL + CWS (Detox<sup>TM</sup>);
- (8) one or more mineral salts (such as an aluminum salt) + a non-toxic derivative of LPS (such as 3dPML).
- (9) one or more mineral salts (such as an aluminum salt) + an immunostimulatory oligonucleotide (such as a nucleotide sequence including a CpG motif). Combination No. (9) is a preferred adjuvant combination.

*M. Human Immunomodulators*

Human immunomodulators suitable for use as adjuvants in the invention include cytokines, such as interleukins (e.g. IL-1, IL-2, IL-4, IL-5, IL-6, IL-7, IL-12, etc.), interferons (e.g. interferon- $\gamma$ ), macrophage colony stimulating factor, and tumor necrosis factor.

Aluminum salts and MF59 are preferred adjuvants for use with injectable influenza vaccines. Bacterial toxins and bioadhesives are preferred adjuvants for use with mucosally-delivered vaccines, such as nasal vaccines.



The immunogenic compositions of the present invention may be administered in combination with an antibiotic treatment regime. In one embodiment, the antibiotic is administered prior to administration of the antigen of the invention or the composition comprising the one or more of the antigens of the invention.

5 In another embodiment, the antibiotic is administered subsequent to the administration of the one or more antigens of the invention or the composition comprising the one or more antigens of the invention. Examples of antibiotics suitable for use in the treatment of the Streptococcal infections of the invention include but are not limited to penicillin or a derivative thereof or clindamycin or the like.

#### 10 Further antigens

The compositions of the invention may further comprise one or more additional Gram positive bacterial antigens which are not associated with an AI. Preferably, the Gram positive bacterial antigens that are not associated with an AI can provide protection across more than one serotype or strain isolate. For example, a first non-AI antigen, in which the first non-AI antigen is at  
 15 least 90% (*i.e.*, at least 90, 91, 92, 93, 94, 95, 96, 97, 98, 99, or 100%) homologous to the amino acid sequence of a second non-AI antigen, wherein the first and the second non-AI antigen are derived from the genomes of different serotypes of a Gram positive bacteria, may be further included in the compositions. The first non-AI antigen may also be homologous to the amino acid sequence of a  
 20 third non-AI antigen, such that the first non-AI antigen, the second non-AI antigen, and the third non-AI antigen are derived from the genomes of different serotypes of a Gram positive bacteria. The first non-AI antigen may also be homologous to the amino acid sequence of a fourth non-AI antigen, such that the first non-AI antigen, the second non-AI antigen, the third non-AI antigen, and the fourth non-AI antigen are derived from the genomes of different serotypes of a Gram positive bacteria.

The first non-AI antigen may be GBS 322. The amino acid sequence of GBS 322 across GBS  
 25 strains from serotypes Ia, Ib, II, III, V, and VIII is greater than 90%. Alternatively, the first non-AI antigen may be GBS 276. The amino acid sequence of GBS 276 across GBS strain from serotypes Ia, Ib, II, III, V, and VIII is greater than 90%. Table 13 provides the percent amino acid sequence identity of GBS 322 and GBS 276 across different GBS strains and serotypes.

Table 13: Conservation of GBS 322 and GBS 276 amino acid sequences

Serotype	Strains	GBS 322		GBS 276	
		cGH	%AA identity	cGH	%AA identity
Ia	090	+	98.60	+	97.90
	A909	+	98.30	+	97.90
	515	+	98.80	+	97.50
	DK1	+		+	
	DK8	+		+	
	Davis	+		+	
Ib	7357b	+		+	
	H36B	+	98.30	+	97.80
II	18RS21	+	100.00	+	99.90
	DK21	+		+	

Serotype	Strains	GBS 322		GBS 276	
		cGH	%AA identity	cGH	%AA identity
III	NEM316	+	100.00	+	97.00
	COH31	+		+	
	D136	+		+	
	M732	+	98.00	+	100.00
	COH1	+	98.30	+	100.00
	M781	+	98.30	+	99.60
No type	CJB110	+	98.60	+	97.90
	1169NT	+	97.40	+	97.90
V	CJB111	+	100.00	+	
	2603	+	100.00	+	100.00
VIII	JM130013	+	100.00	+	97.90
	SMU014	+		+	
total		22/22	98.28+/-0.4	22/22	98.44 +/-1.094

As an example, inclusion of a non-AI protein, GBS 322, in combination with AI antigens GBS 67, GBS 80, and GBS 104 provided protection to newborn mice in an active maternal immunization assay.

5

Table 14: Active maternal immunization assay for a combination of fragments from GBS 322, GBS 80, GBS 104, and GBS 67

GBS strains	Type	FACS ( $\Delta$ Mean)			MIX-322+80+104+67		PBS	
		GBS 80	GBS 67	GBS 322	alive/treated	% protection	alive/treated	% protection
515	Ia	0	409	227	39/40	97	6/40	15
7357b-	Ib	91	316	102	19/30	63	1/30	3
DK21	II	0	331	416	25/34	73	17/48	35
5401	II	170	618	135	35/40	87	3/37	8
3050	II	43	460	188	48/48	100	1/30	3
COH1	III	305	0	130	36/36	100	7/40	17
M781	III	65	0	224	30/40	75	4/39	10
2603	V	125	105	313	27/33	82	10/35	28
CJB111	V	370	481	63	25/28	89	4/46	9
JM9130013	VIII	597	83	143	37/39	95	5/40	12
JMU071	VIII	556	79	170	44/50	88	18/50	36
NT1169	NT	0	443	213	12/32	37	11/35	31

In fact, the non-AI GBS 322 antigen may itself provide protection to newborn mice in an active maternal immunization assay.



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Table 16: Active maternal immunization assay for each of GBS 80 and GBS 322 antigens

GBS strains	Type	GBS 80			GBS 322		
		FACS	Protection (% survival)		FACS	Protection (% survival)	
		$\Delta$ Mean	antigen	ctrl-	$\Delta$ Mean	antigen	ctrl-
CJB111	V	370	72 %	40%	63	57%	40%
COH1	III	305	76 %	10%	130	3%	10%
2603	V	82	22 %	34%	313	83 %	34%
7357b-	Ib	91	36%	34%	102	43%	34%
18RS21	II	0	15%	24%	268	84 %	24%
DK21	II	0	10%	21%	416	67 %	25%
A909	Ia	0	0%	14%			
O90	Ia	0	0%	0%			
H36B	Ib				105	34%	32%

Thus, inclusion of a non-AI protein in an immunogenic composition of the invention may provide increased protection a mammal.

The immunogenic compositions comprising *S. pneumoniae* AI polypeptides may further secondary SP protein antigens which include (a) any of the SP protein antigens disclosed in WO 02/077021 or U.S. provisional application \_\_\_\_\_, filed April 20, 2005 (Attorney Docket Number 002441.00154), (2) immunogenic portions of the antigens comprising at least 7 contiguous amino acids, (3) proteins comprising amino acid sequences which retain immunogenicity and which are at least 95% identical to these SP protein antigens (e.g., 95%, 96%, 97%, 98%, 99%, or 99.5% identical), and (4) fusion proteins, including hybrid SP protein antigens, comprising (1)-(3).

Alternatively, the invention may include an immunogenic composition comprising a first and a second Gram positive bacteria non-AI protein, wherein the polynucleotide sequence encoding the sequence of the first non-AI protein is less than 90% (i.e., less than 90, 88, 86, 84, 82, 81, 78, 76, 74, 72, 70, 65, 60, 55, 50, 45, 40, 35, or 30 percent) homologous than the corresponding sequence in the genome of the second non-AI protein.

The compositions of the invention may further comprise one or more additional non-Gram positive bacterial antigens, including additional bacterial, viral or parasitic antigens. The compositions of the invention may further comprise one or more additional non-GBS antigens, including additional bacterial, viral or parasitic antigens.

In another embodiment, the GBS antigen combinations of the invention are combined with one or more additional, non-GBS antigens suitable for use in a vaccine designed to protect elderly or immunocomprised individuals. For example, the GBS antigen combinations may be combined with an antigen derived from the group consisting of *Enterococcus faecalis*, *Staphylococcus aureus*, *Staphylococcus epidermis*, *Pseudomonas aeruginosa*, *Legionella pneumophila*, *Listeria monocytogenes*, *Neisseria meningitides*, influenza, and Parainfluenza virus ('PIV').

Where a saccharide or carbohydrate antigen is used, it is preferably conjugated to a carrier protein in order to enhance immunogenicity {e.g. Ramsay *et al.* (2001) *Lancet* 357(9251):195-196; Lindberg (1999) *Vaccine* 17 Suppl 2:S28-36; Buttery & Moxon (2000) *J R Coll Physicians Lond* 34:163-168; Ahmad & Chapnick (1999) *Infect Dis Clin North Am* 13:113-133, vii.; Goldblatt (1998) *J. Med. Microbiol.* 47:563-567; European patent 0 477 508; US Patent No. 5,306,492; International patent application WO98/42721; *Conjugate Vaccines* (eds. Cruse *et al.*) ISBN 3805549326, particularly vol. 10:48-114; and Hermanson (1996) *Bioconjugate Techniques* ISBN: 0123423368 or 012342335X}. Preferred carrier proteins are bacterial toxins or toxoids, such as diphtheria or tetanus toxoids. The CRM<sub>197</sub> diphtheria toxoid is particularly preferred {*Research Disclosure*, 453077 (Jan 2002)}. Other carrier polypeptides include the *N.meningitidis* outer membrane protein (EP-A-0372501), synthetic peptides (EP-A-0378881; EP-A-0427347), heat shock proteins (WO 93/17712; WO 94/03208), pertussis proteins (WO 98/58668; EP A 0471177), protein D from *H.influenzae* (WO 00/56360), cytokines (WO 91/01146), lymphokines, hormones, growth factors, toxin A or B from *C.difficile* (WO00/61761), iron-uptake proteins (WO01/72337), *etc.* Where a mixture comprises capsular saccharides from both serogroups A and C, it may be preferred that the ratio (w/w) of MenA saccharide:MenC saccharide is greater than 1 (e.g. 2:1, 3:1, 4:1, 5:1, 10:1 or higher). Different saccharides can be conjugated to the same or different type of carrier protein. Any suitable conjugation reaction can be used, with any suitable linker where necessary.

Toxic protein antigens may be detoxified where necessary e.g. detoxification of pertussis toxin by chemical and/or genetic means.

Where a diphtheria antigen is included in the composition it is preferred also to include tetanus antigen and pertussis antigens. Similarly, where a tetanus antigen is included it is preferred also to include diphtheria and pertussis antigens. Similarly, where a pertussis antigen is included it is preferred also to include diphtheria and tetanus antigens.

Antigens in the composition will typically be present at a concentration of at least 1 µg/ml each. In general, the concentration of any given antigen will be sufficient to elicit an immune response against that antigen.

As an alternative to using protein antigens in the composition of the invention, nucleic acid encoding the antigen may be used {e.g. refs. Robinson & Torres (1997) *Seminars in Immunology* 9:271-283; Donnelly *et al.* (1997) *Annu Rev Immunol* 15:617-648; Scott-Taylor & Dalglish (2000) *Expert Opin Investig Drugs* 9:471-480; Apostolopoulos & Plebanski (2000) *Curr Opin Mol Ther* 2:441-447; Ilan (1999) *Curr Opin Mol Ther* 1:116-120; Dubensky *et al.* (2000) *Mol Med* 6:723-732; Robinson & Pertmer (2000) *Adv Virus Res* 55:1-74; Donnelly *et al.* (2000) *Am J Respir Crit Care Med* 162(4 Pt 2):S190-193; and Davis (1999) *Mt. Sinai J. Med.* 66:84-90}. Protein components of the compositions of the invention may thus be replaced by nucleic acid (preferably DNA e.g. in the form of a plasmid) that encodes the protein.



Definitions

~~PCT/US2005/027239~~

The term "comprising" means "including" as well as "consisting" *e.g.* a composition "comprising" X may consist exclusively of X or may include something additional *e.g.* X + Y.

The term "about" in relation to a numerical value *x* means, for example,  $x \pm 10\%$ .

5       References to a percentage sequence identity between two amino acid sequences means that, when aligned, that percentage of amino acids are the same in comparing the two sequences. This alignment and the percent homology or sequence identity can be determined using software programs known in the art, for example those described in section 7.7.18 of *Current Protocols in Molecular Biology* (F.M. Ausubel *et al.*, eds., 1987) Supplement 30. A preferred alignment is determined by the  
10       Smith-Waterman homology search algorithm using an affine gap search with a gap open penalty of 12 and a gap extension penalty of 2, BLOSUM matrix of 62. The Smith-Waterman homology search algorithm is disclosed in Smith & Waterman (1981) *Adv. Appl. Math.* 2: 482-489.

The invention is further illustrated, without limitation, by the following examples.

**EXAMPLE 1: Binding of an Adhesin Island surface protein, GBS 80, to Fibrinogen and**  
15       **Fibronectin.**

This example demonstrates that an Adhesin Island surface protein, GBS 80 can bind to fibrinogen and fibronectin.

An enzyme-linked immunosorbent assay (ELISA) was used to analyse the *in vitro* binding ability of recombinant GBS 80 to immobilized extra-cellular matrix (ECM) proteins but not to bovine  
20       serum albumin (BSA). Microtiter plates were coated with ECM proteins (fibrinogen, fibronectin, laminin, collagen type IV) and binding assessed by adding varying concentrations of a recombinant form of GBS 80, over-expressed and purified from *E. coli* (FIGURE 5A). Plates were then incubated sequentially with a) mouse anti-GBS 80 primary antibody; b) rabbit anti-mouse AP-conjugated secondary antibody; c) pNPP colorimetric substrate. Relative binding was measured by monitoring  
25       absorbance at 405 nm, using 595 nm as a reference wavelength. Figure 5b shows binding of recombinant GBS 80 to immobilized ECM proteins (1  $\mu$ g) as a function of concentration of GBS 80. BSA was used as a negative control. Data points represent the means of OD<sub>405</sub> values  $\pm$  standard deviation for 3 wells.

Binding of GBS 80 to the tested ECM proteins was found to be concentration dependent and  
30       exhibited saturation kinetics. As is also evident from FIGURE 5, binding of GBS 80 to fibronectin and fibrinogen was greater than binding to laminin and collagen type IV at all the concentrations tested.

**EXAMPLE 2: GBS 80 is required for surface localization of GBS 104.**

This example demonstrates that co-expression of GBS 80 is required for surface localization  
35       of GBS 104.

The polycistronic nature of the Adhesin Island I mRNA was investigated through reverse transcriptase-PCR (RT-PCR) analysis employing primers designed to detect transcripts arising from contiguous genes. Total RNA was isolated from GBS cultures grown to an optical density at 600 nm

(OD<sub>600</sub>) of 0.3 in THB (Todd-Hewitt broth) by the RNeasy Total RNA isolation method (Qiagen) according to the manufacturer's instructions. The absence of contaminating chromosomal DNA was confirmed by failure of the gene amplification reactions to generate a product detectable by agarose gel electrophoresis, in the absence of reverse transcriptase. RT-PCR analysis was performed with the  
5 Access RT-PCR system (Promega) according to the manufacturer's instructions, employing PCR cycling temperatures of 60°C for annealing and 70°C for extension. Amplification products were visualized alongside 100-bp DNA markers in 2% agarose gels after ethidium bromide staining.

FIGURE 5 shows that all the genes are co-transcribed as an operon. A schematic of the AI-1 operon is shown above the agarose gel analysis of the RT-PCR products. Large rectangular arrows  
10 indicate the predicted transcript direction. Primer pairs were selected such as "1-4" cross the 3' finish-5' start of successive genes and overlap each gene by at least 200 bp. Additionally, "1" crosses a putative rho-independent transcriptional terminator. "5" is an internal GBS 80 control and "6" is an unrelated control from a highly expressed gene. Lanes: "a": RNA plus RTase enzyme; "b" RNA without RTase; "c": genomic DNA control.

15 In the effort to elucidate the functions of the AI-1 proteins, in frame deletions of all of the genes within the operon have been constructed and the resulting mutants characterized with respect to surface exposure of the encoded antigens (see FIGURE 8).

Each in-frame deletion mutation was constructed by splice overlap extension PCR (SOE-PCR) essentially as described by Horton et al. [Horton R. M., Z. L. Cai, S. N. Ho, L. R. Pease (1990)  
20 Biotechniques 8:528-35] using suitable primers and cloned into the temperature sensitive shuttle vector pJRS233 to replace the wild type copy by allelic exchange [Perez-Casal, J., J. A. Price, et al. (1993) Mol Microbiol 8(5): 809-19.]. All plasmid constructions utilized standard molecular biology techniques, and the identities of DNA fragments generated by PCR were verified by sequencing. Following SOE-PCR, the resulting mutant DNA fragments were digested with XhoI and EcoRI, and  
25 ligated into a similarly digested pJRS233. The resulting vectors were introduced by electroporation into the chromosome of 2603 and COH1 GBS strains in a three-step process, essentially as described in Framson et al. [Framson, P. E., A. Nittayajarn, J. Merry, P. Youngman, and C. E. Rubens. (1997) Appl. Environ. Microbiol. 63(9):3539-47]. Briefly, the vector pJRS233 contains an *erm* gene encoding erythromycin resistance and a temperature-sensitive gram-positive replicon that is active at  
30 30°C but not at 37°C. Initially, the constructs are electroporated into GBS electro-competent cells prepared as described by Frameson et al., and transformants containing free plasmid are selected by their ability to grow at 30°C on Todd-Hewitt Broth (THB) agar plates containing 1 µg/ml erythromycin. The second step includes a selection step for strains in which the plasmid has integrated into the chromosome via a single recombination event over the homologous plasmid insert and  
35 chromosome sequence by their ability to grow at 37°C on THB agar medium containing 1 mg/ml erythromycin. In the third step, GBS cells containing the plasmid integrated within the chromosome (integrants) are serially passed in broth culture in the absence of antibiotics at 30°C. Plasmid excision



from the chromosome via a second recombination event over the duplicated target gene sequence  
either completed the allelic exchange or reconstituted the wild-type genotype. Subsequent loss of the  
plasmid in the absence of antibiotic selection pressure resulted in an erythromycin-sensitive  
phenotype. In order to assess gene replacement a screening of erythromycin-sensitive colonies was  
performed by analysis of the target gene PCR amplicons.

FIGURE 7 reports a schematic of the IS-1 operon for each knock-out strain generated, along  
with the deletion position within the amino acidic sequence. Most data presented here concern the  
COH1 deletion strains, in which the expression of each of the antigens is higher by DNA microarray  
analysis (data not shown) as well as detectable by FACS analysis (see FIGURE 8). The double mutant  
in 2603  $\Delta 80$ ,  $\Delta 104$  double mutant was constructed by sequential allelic exchanges of the shown  
alleles.

#### *Immunization protocol*

Immune sera for FACS experiments were obtained as follows.

Groups of 4 CD-1 outbred female mice 6-7 weeks old (Charles River Laboratories, Calco  
Italy) were immunized with the selected GBS antigens, (20  $\mu$ g of each recombinant GBS antigen),  
suspended in 100  $\mu$ l of PBS. Each group received 3 doses at days 0, 21 and 35. Immunization was  
performed through intra-peritoneal injection of the protein with an equal volume of Complete  
Freund's Adjuvant (CFA) for the first dose and Incomplete Freund's Adjuvant (IFA) for the following  
two doses. In each immunization scheme negative and positive control groups are used. Immune  
response was monitored by using serum samples taken on day 0 and 49.

#### *FACS analysis*

Preparation of paraformaldehyde treated GBS cells and their FACS analysis were carried out  
as follows.

GBS serotype COH1 strain cells were grown in Todd Hewitt Broth (THB; Difco  
Laboratories, Detroit, Mich.) to OD600nm = 0.5. The culture was centrifuged for 20 minutes at 5000  
rpm and bacteria were washed once with PBS, resuspended in PBS containing 0.05%  
paraformaldehyde, and incubated for 1 hours at 37 °C and then overnight at 4°C. 50 $\mu$ l of fixed  
bacteria (OD600 0.1) were washed once with PBS, resuspended in 20 $\mu$ l of Newborn Calf Serum,  
(Sigma) and incubated for 20 min. at room temperature. The cells were then incubated for 1 hour at  
4°C in 100 $\mu$ l of preimmune or immune sera, diluted 1:200 in dilution buffer (PBS, 20% Newborn  
Calf Serum, 0.1% BSA). After centrifugation and washing with 200 $\mu$ l of washing buffer (0.1% BSA  
in PBS), samples were incubated for 1 hour at 4°C with 50 $\mu$ l of R-Phicoerytrin conjugated F(ab)2  
goat anti-mouse IgG (Jackson ImmunoResearch Laboratories; Inc.), diluted 1:100 in dilution buffer.  
Cells were washed with 200 $\mu$ l of washing buffer and resuspended in 200 $\mu$ l of PBS. Samples were  
analysed using a FACS Calibur apparatus (Becton Dickinson, Mountain View, Calif.) and data were  
analyzed using the Cell Quest Software (Becton Dickinson). A shift in mean fluorescence intensity of  
> 75 channels compared to preimmune sera from the same mice was considered positive. This cutoff

was determined from the mean plus two standard deviations of shifts obtained with control sera raised against mock purified recombinant proteins from cultures of *E. coli* carrying the empty expression vector and included in every experiment. Artifacts due to bacterial lysis were excluded using antisera raised against 6 different known cytoplasmic proteins all of which were negative

5 FACS data on COH1 single KO mutants for GBS 104 and GBS 80 indicated that GBS 80 is required for surface localization of GBS 104.

As shown in FIGURE 8, GBS 104 is not surface exposed in the  $\Delta 80$  strain (second column, bottom), but is present in the whole protein extracts (see FIGURE 10). Mean shift values suggest that GBS 104 is partially responsible for GBS 80 surface exposure (Mean shift of GBS 80 is reduced to  
10  $\sim 60\%$  wild-type levels in  $\Delta 104$ ), and that GBS 80 is over-expressed in the complemented strain (mean shift value  $\sim 200\%$  wild-type level). The  $\Delta 80/p$ GBS 80 strain contains the GBS 80 orf cloned in the shuttle-vector pAM401 (Wirth, R., F. Y. An, et al. (1986). J Bacteriol 165(3): 831-6). The vector alone does not alter the secretion pattern of GBS 104 (right column). FACS was performed on mid-log fixed bacteria with mouse polyclonal antibodies as indicated at left. Black peak is pre-immune  
15 sera, colored peaks are sera from immunized animals.

### EXAMPLE 3: Deletion of GBS 80 causes attenuation *in vivo*.

This example demonstrates that deletion of GBS 80 causes attenuation *in vivo*, suggesting that this protein contributes to bacterial virulence.

20 By using a mouse animal model, we studied the role of GBS 80 and GBS 104 in the virulence of *S. agalactiae*.

Groups of ten outbred female mice 5-6 week weeks old (Charles River Laboratories, Calco Italy) were inoculated intraperitoneally with different dilutions of the mutant strains and LD<sub>50</sub> (lethal dose 50) were calculated according to the method of Reed and Muench [Reed, L. J. and H. Muench  
25 (1938).The American Journal of Hygiene 27(3): 493-7]. As presented in the table below the number of colony forming units (cfu) counted for both the  $\Delta 80$  and the  $\Delta 80, \Delta 104$  double mutants is about 10 fold higher when compared to the wild type strain suggesting that inactivation of GBS 80 but not GBS 104 is responsible for an attenuation in virulence. This finding indicates that GBS 80 gene in the AI-1 might contribute to virulence.

30 Table Lethal dose 50% analysis of AI-1 mutants in the 2603 strain background. LD<sub>50</sub>s were performed by IP injection of female CD1 mice at an age of 5-6 weeks. LD<sub>50</sub>s were calculated by the method of Reed and Muench (8).

GBS strain	LD <sub>50</sub> , cfu	Number of Experiments
Wild Type 2603	$2 \times 10^8$	4
$\Delta 104$ mutant	$\sim 2 \times 10^8$	1
$\Delta 80$ mutant	$2.6 \times 10^9$	3
$\Delta 80, \Delta 104$ double mutant	$\sim 2 \times 10^9$	1

### EXAMPLE 4: Effect of Adhesin Island Sortase Deletions on Surface Antigen Presentation



This example demonstrates the effect of adhesin island sortase deletions on surface antigen presentation.

FACS analysis results set forth in FIGURE 9 show that a deletion in sortase SAG0648 prevented GBS 104 from reaching the surface and slightly reduced the surface exposure of GBS 80 (fourth panel; mean shift value ~60% wild-type COH1). In the double sortase knock-out strain, neither antigen was surface exposed (far right panel). Either sortase alone was sufficient for GBS 80 to arrive at the bacterial surface (third and fourth columns, top). No effect was seen on surface exposure of antigens GBS 80 or GBS 104 in the  $\Delta$ GBS 52 strain. Antibodies derived from purified GBS 52 were either non-specific or were FACS negative for GBS 52 (data not shown). FACS analysis was performed as described above (see EXAMPLE 2).

As shown in FIGURE 10, inactivation of GBS 80 has no effect on GBS 104 expression as much as GBS 104 knock out doesn't change the total amount GBS 80 expressed. The Western blot of whole protein extracts (strains noted above lanes) probed with anti-GBS 80 antisera is shown in panel A. Arrow indicates expected size of GBS 80 (60 kDa). GBS 80 antibodies recognize a doublet, the lower band is not present in  $\Delta$ GBS 80 strains. Panel B shows a Western blot of whole protein extracts probed with anti-GBS 104 antisera. Arrow indicates expected size of GBS 104 (99.4 kDa). Protein extracts were prepared from the same bacterial cultures used for FACS (FIGURES 8 and 9). In conclusion, although GBS 104 does not arrive at the surface in the  $\Delta$ 80 strain by FACS (FIGURE 8, second column), it is present at approximately wild-type levels in the whole protein preps (B, second lane). Approximately 20  $\mu$ g of each protein extract was loaded per lane.

#### *Western-blot analysis*

Aliquots of total protein extract mixed with SDS loading buffer (1x: 60 mM TRIS-HCl pH 6.8, 5% w/v SDS, 10% v/v glycerol, 0.1% Bromophenol Blue, 100 mM DTT) and boiled 5 minutes at 95° C, were loaded on a 12.5% SDS-PAGE precast gel (Biorad). The gel is run using a SDS-PAGE running buffer containing 250 mM TRIS, 2.5 mM Glycine and 0.1 %SDS. The gel is electroblotted onto nitrocellulose membrane at 200 mA for 60 minutes. The membrane is blocked for 60 minutes with PBS/0.05 % Tween-20 (Sigma), 10% skimmed milk powder and incubated O/N at 4° C with PBS/0.05 % Tween 20, 1% skimmed milk powder, with the appropriate dilution of the sera. After washing twice with PBS/0.05 % Tween, the membrane is incubated for 2 hours with peroxidase-conjugated secondary anti-mouse antibody (Amersham) diluted 1:4000. The nitrocellulose is washed three times for 10 minutes with PBS/0.05 % Tween and once with PBS and thereafter developed by Opti-4CN Substrate Kit (Biorad).

#### **Example 5: Binding of Adhesin Island proteins to epithelial cells and effect of Adhesin Island proteins on capacity of GBS to adhere to epithelial cells.**

This example illustrates the binding of AI proteins to epithelial cells and the effect of AI proteins on the capacity of GBS to adhere to epithelial cells.

Applicants analysed whether recombinant AI surface proteins GBS 80 or GBS 104 would demonstrate binding to various epithelial cells in a FACS analysis. Applicants also analysed whether

deletion of AI surface proteins GBS 80 or GBS 104 would effect the capacity of GBS to adhere to and invade ME180 cervical epithelial cells.

As shown in Figure 28, deletion of GBS 80 sequence from GBS strain isolate 2603 (serotype V) did not affect the capacity of the mutated GBS to adhere to and invade ME180 cervical epithelial cells. Here ME180 cervical carcinoma epithelial cells were infected with wild type GBS 2603 or GBS 2603  $\Delta$ 80 isogenic mutant. After two hours of infection, non-adherent bacteria were washed off and infection prolonged for a further two hours and four hours. In invasion experiments, after each time point, was followed by a two hour antibiotic treatment. Cells were then lysed with 1% saponin and lysates plated on TSA plates. As shown in Figure 28, there was little difference between the percent invasion or percent adhesion of wild type and mutant strains up to the four hour time point.

Figure 30 repeats this experiment with both  $\Delta$ 104 and  $\Delta$  80 mutants from a different strain isolate. Here, ME180 cervical carcinoma epithelial cells were infected with GBS strain isolate COH (serotype III) wild type or COH1  $\Delta$ GBS 104 or COH1  $\Delta$ 80 isogenic mutant. After one hour of infection, non-adherent bacteria were washed off and the cells were lysed with 1% saponin. The lysates were plated on TSA plates. As shown in Figure 30, while there was little difference in the percent invasion, there was a significant decrease in the percent association of the  $\Delta$ 104 mutant compared to both the wild type and  $\Delta$ 80 mutant.

The affect of AI surface proteins on the ability of GBS to translocate through an epithelial monolayer was also analysed. As shown in Figure 31, a GBS 80 knockout mutant strain partially loses the ability to translocate through an epithelial monolayer. Here epithelial monolayers were inoculated with wildtype or knockout mutant in the apical chamber of a transwell system for two hours and then non-adherent bacteria were washed off. Infection was prolonged for a further two and four hours. Samples were taken from the media of the basolateral side and the number of colony forming unities measured. Transepithelial electrical resistance measured prior to and after infection gave comparable values, indicating the maintenance of the integrity of the monolayer. By the six hour time point, the  $\Delta$ 80 mutants demonstrated a reduced percent transcytosis.

A similar experiment was conducted with GBS 104 knock out mutants. Here, as shown in Figure 22, the  $\Delta$ 104 mutants also demonstrated a reduced percent transcytosis, indicating that the mutant strains translocate through an epithelial monolayer less efficiently than their isogenic wild type counterparts.

Applicants also studied the effect of AI proteins on the capacity of a GBS strain to invade J774 macrophage-like cells. Here, J774 cells were infected with GBS COH1 wild type or COH1  $\Delta$ GBS104 or COH1  $\Delta$ GBS80 isogenic mutants. After one hour of infection, non-adherent bacteria were washed off and intracellular bacteria were recovered at two, four and six hours post antibiotic treatment. At each time point, cells were lysed with 0.25% Triton X-100 and lysates plated on TSA plates. As shown in Figure 32, the  $\Delta$ 104 mutant demonstrated a significantly reduced percent invasion compared to both the wild type and  $\Delta$ 80 mutant.



**Example 6: Hyperoligomeric structures comprising AI surface proteins GBS 80 and GBS 104.**

This example illustrates hyperoligomeric structures comprising AI surface proteins GBS 80 and GBS 104. A GBS isolate COH1 (serotype III) was adapted to increase expression of GBS 80. Figure 34 presents a regular negative stain electron micrograph of this mutant; no pilus or hyperoligomeric structures are distinguishable on the surface of the bacteria. When the EM stain is based on anti-GBS 80 antibodies labelled with 10 or 20 nm gold particles, the presence of GBS 80 throughout the hyperoligomeric structure is clearly indicated (Figures 36, 37 and 38). EM staining against GBS 104 (anti-GBS 104 antibodies labelled with 10 nm gold particles) also reveals the presence of GBS 104 primarily on or near the surface of the bacteria or potentially associated with bacterial peptidoglycans (Figure 39). Analysis of this same strain (over-expressing GBS 80) with a combination of both anti-GBS 80 (using 20 nm gold particles) and anti-GBS 104 (using 10 nm gold particles) reveals the presence of GBS 104 on the surface and within the hyperoligomeric structures (see Figures 40 and 41).

**Example 7: GBS 80 is necessary for polymer formation and GBS 104 and sortase SAG0648 are necessary for efficient pili assembly**

This example demonstrates that GBS 80 is necessary for formation of polymers and that GBS 104 and sortase SAG0648 are necessary for efficient pili assembly. GBS 80 and GBS 104 polymeric assembly was systematically analyzed in Coh1 strain single knock out mutants of each of the relevant coding genes in AI-1 (GBS 80, GBS 104, GBS 52, sag0647, and sag0648). Figure 41 provides Western blots of total protein extracts (strains noted above lanes) probed with either anti-GBS 80 (left panel) sera or anti-GBS 104 sera (right panel) for each of these Coh1 and Coh1 knock out strains. (Coh1, wild type Coh1;  $\Delta$ 80, Coh1 with GBS 80 knocked out;  $\Delta$ 104, Coh1 with GBS 104 knocked out;  $\Delta$ 52, Coh1 with GBS 52 knocked out;  $\Delta$ 647, Coh1 with SAG0647 knocked out;  $\Delta$ 648, Coh1 with SAG0648 knocked out;  $\Delta$ 647-8, Coh1 with SAG0647 and SAG0648 knocked out;  $\Delta$ 80/pGBS80, Coh1 with GBS 80 knocked out but complemented with a high copy number plasmid expressing GBS 80. Asterisks identify the monomer of GBS 80 and GBS 104.)

The smear of immunoreactive material observed in the wild type strain, along with its disappearance in  $\Delta$ 80 and  $\Delta$ 104 mutants, is consistent with the notion that such high molecular weight structures are composed of covalently linked (SDS-resistant) GBS 80 and GBS 104 subunits. The immunoblotting with both anti-GBS 80 ( $\alpha$ -GBS 80) and anti-GBS 104 ( $\alpha$ -GBS 104) revealed that deletion of sortase SAG0648 also interferes with the assembly of high molecular weight species, whereas the knock out mutant of the second sortase (SAG0647), even if somehow reduced, still maintains the ability to form polymeric structures.

Total extracts from GBS were prepared as follows. Bacteria were grown in 50 ml of Todd-Hewitt broth (Difco) to an OD<sub>600nm</sub> of 0.5-0.6 and successively pelleted. After two washes in PBS the pellet was resuspended and incubated 3 hours at 37°C with mutanolisin. Cells were then lysed with at

least three freezing-thawing cycles in dry ice and a 37°C bath. The lysate was then centrifuged to eliminate the cellular debris and the supernatant was quantified. Approximately 40 µg of each protein extract was separated on SDS-PAGE. The gel was then subjected to immunoblotting with mice antisera and detected with chemiluminescence.

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**Example 8: GBS 80 is polymerized by an AI-2 sortase**

This example illustrates that GBS 80 can be polymerized not only by AI-1 sortases, but also by AI-2 sortases. Figure 42 shows total cell extract immunoblots of GBS 515 strain, which lacks AI-1. The left panel, where an anti-GBS 67 sera was used, shows that GBS 67 from AI-2 is assembled into high-molecular weight-complexes, suggesting the formation of a second type of pilus. The same high molecular structure is observed when GBS 80 is highly expressed by reintroducing the gene within a plasmid (pGBS 80). By using anti-GBS 80 (right panel) sera on the same extracts, again it is observed that, with GBS 80 over expression (515/pGBS 80), a high-molecular weight structure is assembled. This implies that, in the absence of AI-1 sortases, AI-2 sortases (SAG1405 and SAG1406) can complement the lacking function, still being able to assemble GBS 80 in a pilus structure.

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**Example 9: Coh1 produces a high molecular weight molecule, the GBS 80 pilin**

This example illustrates that Coh1 produces a high molecular weight molecule, greater than 1000 kDa, which is the GBS 80 pilin. Figure 43 provides silver-stained electrophoretic gels that show that Coh1 produces two macromolecules. One of these macromolecules disappears in the Coh1 GBS 80 knock out cells, but does not disappear in the Coh1 GBS 52 knock out mutant cells. The last two lanes on the right were loaded with 15 times the amount loaded in the other lanes. This was done in order to be able to count the bands. By doing this, a conservative size estimate of the top bands was calculated by starting at 240 kDa and considering each of 14 higher bands as the result of consecutive additions of a GBS 80 monomer.

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Coh1, wild type Coh1; Δ80, Coh1 cells with GBS 80 knocked out; Δ52, Coh1 cells with GBS 52 knocked out; Δ80/pGBS 80, Coh1 cells with GBS 80 knocked out and complemented with a high copy number construct expressing GBS 80.

**Example 10. GBS 52 is a minor component of the GBS pilus**

This example illustrates that GBS 52 is present in the GBS pilus and is a minor component of the pilus. Figure 45 shows an immunoblot of total cell extracts from a GBS Coh1 strain and a GBS Coh1 strain knocked out for GBS 52 (Δ52). The total cell extracts were immunoblotted anti-GBS 80 antisera (left) and anti-GBS 52 antisera (right). Immunoblotting was performed using a 3-8% Tris-acetate polyacrylamide gel (Invitrogen) which provided excellent separation of large molecular weight proteins (see figure 41). When the gel was incubated with anti-GBS 80 sera, the bands from the Coh1 wild-type strain appeared shifted when compared to the Δ52 mutant. This observation

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indicated a different size of the pilus polymeric components in the two strains. When the same gel was stripped and incubated with anti-GBS 52 sera the high-molecular subunits in the Coh1 wild-type strain showed similar molecular size of those in the correspondent lane in the left panel. These findings confirmed that GBS 52 is indeed associated with GBS 80 macro-molecular structures but represents a minor component of the GBS pilus.

#### **Example 11: Pilus structures are present in the supernatant of GBS bacterial cultures**

This example illustrates that the pilus structure assembled in Coh1 GBS is present in the supernatant of a bacterial cell culture. Figure 46 shows an immunoblot where the protein extract of the supernatant from cultures of different GBS mutant strains (117 = Coh1 GBS 80 knockout; 159 = Coh1 GBS 104 knockout; 202 = Coh1 GBS 52 knockout; 206 = Coh1 GBS sag0647 knockout; 208 = Coh1 GBS sag0648 knockout; 197 = Coh1 GBS sag0647/sag0648 knockout; 179 = Coh1 GBS 80 knockout complemented with a high copy plasmid expressing GBS 80). GBS 80 antisera detects the presence of pilus structures in the appropriate Coh1 strains.

The protein extract was prepared as follows. Bacteria were grown in THB to an OD<sub>600nm</sub> of 0.5-0.6 and the supernatant was separated from the cells by centrifugation. The supernatant was then filtered ( $\varnothing$  0.2  $\mu$ m) and 1 ml was added with 60% TCA for protein precipitation. GBS pili were also extracted from the fraction of surface-exposed proteins in Coh1 strain and its GBS 80 knock out mutant as described hereafter. Bacteria were grown to an OD<sub>600nm</sub> of 0.6 in 50 ml of THB at 37°C. Cells were washed once with PBS and the pellet was then resuspended in 0.1 M KPO4 pH 6.2, 40% sucrose, 10 mM MgCl<sub>2</sub>, 400U/ml mutanolysin and incubated 3 hours at 37°C. Protoplasts were separated by centrifugation and the supernatant was recovered and its protein content measured.

In order to study the dynamics of pilus production during different growth phases, 1 ml supernatant of a culture at different OD<sub>600nm</sub> was TCA precipitated and loaded onto a 3-8% SDS-PAGE as described before. Figure 47 shows the corresponding Western blot with GBS 80 anti-sera. The first group of lanes (left five sample lanes) refer to a Coh1 strain growth (OD<sub>600nm</sub> are noted above the lanes) whereas the second group of lanes (right five samples) are from a GBS 80 knock out strain over expressing GBS 80. The experiment shows that pilus macromolecular structures can be found in the supernatant in all of the growth phases tested.

#### **Example 12: In GBS strain Coh1, only GBS 80 and a sortase (sag0647 or sag0648) is required for polymerization**

This example describes requirements for pilus formation in Coh1. Figure 48 shows a Western blot of total protein extracts (prepared as described before) using anti-GBS 80 sera on Coh1 clones. (Coh1, wild type Coh1;  $\Delta$ 104, Coh1 knocked out for GBS 104,  $\Delta$ 647, Coh1 knocked out for sag0647,  $\Delta$ 648, Coh1 knocked for sag0648,  $\Delta$ 647-8, Coh1 knocked out for sag0647 and sag0648; 515, wild

type bacterial strain 515, which lacks an AI-1; p80 a high copy number plasmid which expresses GBS 80.) The data show that only the double sortase mutant is unable to polymerize GBS 80 indicating that the 'conditio sine qua non' for pilus polymerization is the co-existence of GBS 80 with at least one sortase. This result leads to a reasonable assumption that SAG1405 and SAG1406 are responsible for polymerization in this strain.

**Example 13: GBS 80 can be expressed in *L. lactis* under its own promoter and terminator sequences**

This example demonstrates that *L. lactis*, a non-pathogenic bacterium, can express GBS AI polypeptides such as GBS 80. *L. lactis* M1363 (*J. Bacteriol.* 154 (1983):1-9) was transformed with a construct encoding GBS 80. Briefly, the construct was prepared by cloning a DNA fragment containing the gene coding for GBS 80 under its own promoter and terminator sequences into plasmid pAM401 (a shuttle vector for *E. coli* and other Gram positive bacteria; *J. Bacteriol.* 163 (1986):831-836). Total extracts of the transformed bacteria in log phase were separated on SDS-PAGE, transferred to membranes, and incubated with antiserum against GBS 80. A polypeptide corresponding to the molecular weight of GBS 80 was detected in the lanes containing total extracts of *L. lactis* transformed with the GBS 80 construct. See Figures 133A and 133B, lanes 6 and 7. This same polypeptide was not detected in the lane containing total extracts of *L. lactis* not transformed with the GBS 80 construct, lane 9. This example shows that *L. lactis* can express GBS 80 under its own promoter and terminator.

**Example 14: *L. lactis* modified to express GBS AI-1 under the GBS 80 promoter and terminator sequences expresses GBS 80 in polymeric structures**

This example demonstrates the ability of *L. lactis* to express GBS AI-1 polypeptides and to incorporate at least some of the polypeptides into oligomers. *L. lactis* was transformed with a construct containing the genes encoding GBS AI-1 polypeptides. Briefly, the construct was prepared by cloning a DNA fragment containing the genes for GBS 80, GBS 52, SAG0647, SAG0648, and GBS 104 under the GBS 80 promoter and terminator sequences into construct pAM401. The construct was transformed into *L. lactis* M1363. Total extracts of log phase transformed bacteria were separated on reducing SDS-PAGE, transferred to membranes, and incubated with antiserum against GBS 80. A polypeptide with a molecular weight corresponding to the molecular weight of GBS 80 was detected in the lanes containing *L. lactis* transformed with the GBS AI-1 encoding construct. See Figure 134, lane 2. In addition, the same lane also showed immunoreactivity of polypeptides having higher molecular weights than the polypeptide having the molecular weight of GBS 80. These higher molecular weight polypeptides are likely oligomers of GBS 80. Oligomers of similar molecular



weights were also observed on a Western blot of the culture supernatant of the transformed *L. lactis*. See lane 4 of Figure 135. Thus, this example shows that *L. lactis* transformed to express GBS AI-1 can efficiently polymerize GBS 80 in the form of a pilus. This pilus structure can likely be purified from either the cell culture supernatant or cell extracts.

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#### Example 15: Cloning and Expression of *S. pneumoniae* Sp0462

This example describes the production of a clone encoding a Sp0462 polypeptide and expression of the clone. To produce a clone encoding Sp0462, the open reading frame encoding Sp0462 was amplified using primers that annealed within the full-length Sp0462 open reading frame sequence. Figure 150A provides a 893 amino acid sequence of Sp0462. The primers used to produce a clone encoding the Sp0462 polypeptide are shown in Figure 150B. These primers annealed to the nucleotide sequences encoding the amino acid residues indicated by underlining in Figure 150A. Amplification of the open reading frame encoding Sp0462 using these primers produced the amplicon shown at lane 2 of the agarose gel provided in Figure 160. The Sp0462 clone encodes amino acid residues 38-862 of the 893 amino acid residue Sp0462 protein; the italicized residues in Figure 150A were eliminated. Figure 151A provides a schematic depiction of the recombinant Sp0462 polypeptide. Figure 151B shows a schematic depiction of the full-length Sp0462 polypeptide. Both the recombinant Sp0462 encoded by the clone and the full-length Sp0462 protein have two collagen binding protein type B (Cna B) domains and a von Hillebrand factor A (vWA) domain. The cloned recombinant Sp0462 lacks the LPXTG motif present in the full-length Sp0462 protein. Western blot analysis for expression of the Sp0462 clone did not result in detection of polypeptides with serum obtained from *S. pneumoniae*-infected patients (Figure 152A) or GBS 80 antiserum (Figure 152B).

#### Example 16: Cloning and Expression of *S. pneumoniae* Sp0463

This example describes the production of a clone encoding a Sp0463 polypeptide and detection of recombinant Sp0463 polypeptide expressed from the clone. To produce a clone encoding Sp0463, the open reading frame encoding Sp0463 was amplified using primers that annealed within the full-length Sp0463 open reading frame sequence. Figure 153A provides a 665 amino acid sequence of Sp0463. The primers used to produce the clone encoding Sp0463 polypeptide are shown in Figure 153B. These primers annealed to the nucleotide sequences encoding the amino acid residues indicated by underlining in Figure 153A. Amplification of the open reading frame encoding Sp0463 using these primers produced the amplicon shown at lane 3 of the agarose gel provided in Figure 160. The Sp0463 clone encodes amino acid residues 23-627 of the 665 amino acid residue Sp0463 protein; the italicized residues in Figure 153A were eliminated. Figure 154A provides a schematic depiction of the recombinant Sp0463 polypeptide. Figure 154B shows a schematic depiction of the full-length Sp0463 polypeptide. Both the recombinant Sp0463 encoded by the clone and the full-length Sp0463 protein have a Cna B domain and an E box motif. The cloned recombinant

Sp0463 lacks the LPXTG motif present in the full-length Sp0463 protein. Expression of the Sp0463 clone resulted in the detection of a 60 kD polypeptide, the expected molecular weight of the recombinant Sp0463 polypeptide, by Western blot analysis. See Figure 155.

5 **Example 17: Cloning and Expression of *S. pneumoniae* Sp0464**

This example describes the production of a clone encoding a Sp0464 polypeptide and detection of recombinant Sp0464 polypeptide expressed from the clone. To produce a clone encoding Sp0464, the open reading frame encoding Sp0464 was amplified using primers that annealed either within the full-length Sp0464 open reading frame sequence. Figure 157A provides a 393 amino acid  
10 sequence of Sp0464. The primers used to produce a clone encoding the Sp0464 polypeptide are shown in Figure 157B. These primers annealed to the nucleotide sequences encoding the amino acid residues indicated by underlining in Figure 157A. Amplification of the open reading frame encoding Sp0464 using these primers produced the amplicon shown at lane 4 of the agarose gel provided in Figure 160. The Sp0464 clone encodes amino acid residues 19-356 of the 393 amino acid residue  
15 Sp0464 protein; the italicized residues in Figure 157A were eliminated. Figure 158A provides a schematic depiction of the recombinant Sp0464 polypeptide. Figure 158B shows a schematic depiction of the full-length Sp0464 polypeptide. Both the recombinant Sp0464 encoded by the clone and the full-length Sp0464 protein have two Cna B domains. The cloned recombinant Sp0464 lacks the LPXTG motif present in the full-length Sp0464 protein. Expression of the Sp0464 clone resulted  
20 in the detection of a 38 kD polypeptide, the expected molecular weight of the recombinant Sp0464 polypeptide, by Western blot analysis. See Figure 159.

**Example 18: Intranasal Immunization of Mice with Recombinant *L. lactis* Expressing GBS 80 and Subsequent Challenge**

This example describes a method of intranasally immunizing mice using *L. lactis* that express GBS 80. Intranasal immunization consisted of 3 doses at days 0, 14 and 28, each dose administered in three consecutive days. Each day, groups of 3 CD-1 outbred female mice 6-7 weeks old (Charles River Laboratories, Calco Italy) were immunized intranasally with  $10^9$  or  $10^{10}$  CFU of the recombinant *Lactococcus lactis* suspended in 20  $\mu$ l of PBS. In each immunization scheme negative  
30 (wild-type *L. lactis*) and positive (recombinant GBS80) control groups were used. The immune response of the dams was monitored by using serum samples taken on day 0 and 49. The female mice were bred 2-7 days after the last immunization (at approximately  $t = 36 - 37$ ), and typically had a gestation period of 21 days. Within 48 hours of birth, the pups were challenged via I.P. with GBS in a dose approximately equal to an amount which would be sufficient to kill 90 % of immunized pups (as  
35 determined by empirical data gathered from PBS control groups). The GBS challenge dose is preferably administered in 50ml of THB medium. Preferably, the pup challenge takes place at 56 to 61 days after the first immunization. The challenge inocula were prepared starting from frozen



cultures diluted to the appropriate concentration with THB prior to use. Survival of pups was monitored for 5 days after challenge.

#### Example 19: Subcutaneous Immunization of Mice with Recombinant *L. lactis* Expressing GBS

##### 5 80 and Subsequent Challenge

This example describes a method of subcutaneous immunization mice using *L. lactis* that express GBS 80. Subcutaneous immunization consists of 3 doses at days 0, 14 and 28. Groups of 3 CD-1 outbred female mice 6-7 weeks old (Charles River Laboratories, Calco Italy) were injected subcutaneously with  $10^9$  or  $10^{10}$  CFU of the recombinant *Lactococcus lactis* suspended in 100  $\mu$ l of  
10 PBS. In each immunization scheme, negative (wild-type *L. lactis*) and positive (recombinant GBS80) control groups were used. The immune response of the dams was monitored by using serum samples taken on day 0 and 49. The female mice were bred 2-7 days after the last immunization (at approximately  $t = 36 - 37$ ), and typically had a gestation period of 21 days. Within 48 hours of birth, the pups were challenged via I.P. with GBS in a dose approximately equal to an amount which would  
15 be sufficient to kill 90 % of immunized pups (as determined by empirical data gathered from PBS control groups). The GBS challenge dose is preferably administered in 50ml of THB medium. Preferably, the pup challenge takes place at 56 to 61 days after the first immunization. The challenge inocula were prepared starting from frozen cultures diluted to the appropriate concentration with THB prior to use. Survival of pups was monitored for 5 days after challenge.

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#### Example 20: Immunization of Mice with GAS AI polypeptides and Subsequent Intranasal Challenge

This example describes a method of immunizing mice with GAS AI polypeptides and subsequently intranasally challenging the mice with GAS bacteria. Groups of 10 CD1 female mice  
25 aged between 6 and 7 weeks are immunized with a combination of GAS antigens of the invention GAS 15, GAS 16, and GAS 18, (15  $\mu$ g of each recombinant antigen, derived from M1 strain SF370) or *L. lactis* expressing the M1 strain SF370 adhesin island, suspended in 100  $\mu$ l of suitable solution. Each group receives 3 doses at days 0, 21 and 45. Immunization is performed through subcutaneous or intraperitoneal injection for the GAS 15, GAS 16, GAS 18 protein combination. The protein  
30 combination is administered with an equal volume of Complete Freund's Adjuvant (CFA) for the first dose and Incomplete Freund's Adjuvant (IFA) for the following two doses. Immunization is performed intranasally for the *L. lactis* expressing the M1 strain SF370 adhesin island. In each immunization scheme negative and positive control groups are used.

The negative control group for the mice immunized with the GAS 15, GAS 16, GAS 18  
35 protein combination included mice immunized with PBS. The negative control group for the mice immunized with *L. lactis* expressing the M1 strain SF370 adhesin island, included mice immunized

with either wildtype *L. lactis* or *L. lactis* transformed with the pAM401 expression vector lacking any cloned adhesin island sequence.

The positive control groups included mice immunized with purified M1 strain SF370 M protein.

- 5 Immunized mice are then anaesthetized with Zoletil and challenged intranasally with a 25  $\mu$ L suspension containing  $1.2 \times 10^6$  or  $1.2 \times 10^8$  CFU of ISS 3348 in THB. Animals are observed daily and checked for survival.

#### Example 21: Active Maternal Immunization Assay

- 10 As used herein, an Active Maternal Immunization assay refers to an *in vivo* protection assay where female mice are immunized with the test antigen composition. The female mice are then bred and their pups are challenged with a lethal dose of GBS. Serum titers of the female mice during the immunization schedule are measured as well as the survival time of the pups after challenge.

#### 15 Mouse immunization

- Specifically, groups of 4 CD-1 outbred female mice 6-8 weeks old (Charles River Laboratories, Calco Italy) are immunized with one or more GBS antigens, (20  $\mu$ g of each recombinant GBS antigen), suspended in 100  $\mu$ L of PBS. Each group receives 3 doses at days 0, 21 and 35. Immunization is performed through intra-peritoneal injection of the protein with an equal volume of Complete Freund's Adjuvant (CFA) for the first dose and Incomplete Freund's Adjuvant (IFA) for the following two doses. In each immunization scheme negative and positive control groups are used.

Immune response is monitored by using serum samples taken on day 0 and 49. The sera are analyzed as pools from each group of mice.

#### 25 Active maternal immunization

- A maternal immunization/neonatal pup challenge model of GBS infection was used to verify the protective efficacy of the antigens in mice. The mouse protection study was adapted from Rodewald et al. (Rodewald et al. J. Infect. Diseases 166, 635 (1992)). In brief, CD-1 female mice (6-8 weeks old) were immunized before breeding, as described above. The mice received 20  $\mu$ g of protein per dose when immunized with a single antigen and 60  $\mu$ g of protein per dose (15  $\mu$ g of each antigen) when immunized with the combination of antigens. Mice were bred 2-7 days after the last immunization. Within 48 h of birth, pups were injected intraperitoneally with 50  $\mu$ L of GBS culture. Challenge inocula were prepared starting from frozen cultures diluted to the appropriate concentration with THB before use. In preliminary experiments (not shown), the challenge doses per pup for each strain tested were determined to cause 90% lethality. Survival of pups was monitored for 2 days after challenge. Protection was calculated as (percentage



$$\frac{\text{percentage dead Vaccine} - \text{percentage dead Control}}{\text{percentage dead Control}} \times 100$$
 divided by percentage dead Control multiplied by 100. Data were evaluated for statistical significance by Fisher's exact test.

Embodiments of the Invention

The invention encompasses, but is not limited to, the embodiments enumerated below.

1. An immunogenic composition comprising a purified Group B Streptococcus (GBS) adhesin island (AI) polypeptide in oligomeric form.

2. The immunogenic composition of embodiment 1 wherein the GBS AI polypeptide is selected from a GBS AI-1.

3. The immunogenic composition of embodiment 1 wherein the GBS AI polypeptide is selected from a GBS AI-2.

1. An immunogenic composition comprising a purified Group B Streptococcus (GBS) adhesin island (AI) polypeptide in oligomeric form.

2. The immunogenic composition of embodiment 1 wherein the GBS AI polypeptide is selected from a GBS AI-1.

3. The immunogenic composition of embodiment 1 wherein the GBS AI polypeptide is selected from a GBS AI-2.

4. The immunogenic composition of any of embodiments 1-3 wherein the GBS AI polypeptide comprises a sortase substrate motif.

5. The immunogenic composition of embodiment 4 wherein the sortase substrate motif is an LPXTG motif.

6. The immunogenic composition of embodiment 5 wherein the LPXTG motif is represented by the amino acid sequence XPXTG, wherein the X at amino acid position 1 is an L, an I, or an F and the X at amino acid position 3 is any amino acid residue.

7. The immunogenic composition of any one of embodiments 1-3 wherein the GBS AI polypeptide affects the ability of GBS bacteria to adhere to epithelial cells.

8. The immunogenic composition of any one of embodiments 1-3 wherein the GBS AI polypeptide affects the ability of GBS bacteria to invade epithelial cells.

9. The immunogenic composition of any one of embodiments 1-3 wherein the GBS AI polypeptide affects the ability of GBS bacteria to translocate through an epithelial cell layer.

10. The immunogenic composition of any one of embodiments 1-3 wherein the GBS AI polypeptide is capable of associating with an epithelial cell surface.

11. The immunogenic composition of embodiment 10 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.

12. The immunogenic composition of any of embodiments 1-3 wherein the GBS AI polypeptide is a full-length GBS AI protein.

13. The immunogenic composition of any of embodiments 1-3 wherein the GBS AI polypeptide is a fragment of a full-length GBS AI protein.

14. The immunogenic composition of embodiment 13 wherein the fragment comprises at least 7 contiguous amino acid residues of the GBS AI protein.



15. The immunogenic composition of embodiment 2 wherein the GBS AI polypeptide is selected from the group consisting of GBS 80, GBS 104, GBS 52, and fragments thereof.

16. The immunogenic composition of embodiment 3 wherein the GBS AI polypeptide is selected from the group consisting of GBS 59, GBS 67, GBS 150, 01521, 01523, 01524, and fragments thereof.

17. The immunogenic composition of embodiment 15 wherein the GBS AI polypeptide is GBS 80.

18. The immunogenic composition of any of embodiments 1-3 or 15-17 wherein the oligomeric form is a hyperoligomer.

19. The immunogenic composition of any of embodiments 1-3, or 15-17 further comprising a Gram positive bacterium antigen not associated with an AI.

20. The immunogenic composition of embodiment 19 wherein the antigen is selected from the group consisting of GBS 322 and GBS 276.

21. The immunogenic composition of embodiment 20 wherein the antigen is GBS 322.

22. An immunogenic composition comprising a purified Gram positive bacteria adhesin island (AI) polypeptide in an oligomeric form.

23. The immunogenic composition of embodiment 22 wherein the Gram positive bacteria is of a genus selected from the group consisting of *Streptococcus*, *Enterococcus*, *Staphylococcus*, or *Listeria*.

24. The immunogenic composition of embodiment 23 wherein the Gram positive bacteria is of the genus *Streptococcus*.

25. The immunogenic composition of any of embodiments 22-24 wherein the Gram positive bacteria AI polypeptide comprises a sortase substrate motif.

26. The immunogenic composition of embodiment 25 wherein the sortase substrate motif is an LPXTG motif.

27. The immunogenic composition of any one of embodiments 22-24 wherein the Gram positive bacteria AI polypeptide affects the ability of Gram positive bacteria to adhere to epithelial cells.

28. The immunogenic composition of any one of embodiments 22-24 wherein the Gram positive bacteria AI polypeptide affects the ability of Gram positive bacteria to invade epithelial cells.

29. The immunogenic composition of any one of embodiments 22-24 wherein the Gram positive bacteria AI polypeptide affects the ability of Gram positive bacteria to translocate through an epithelial cell layer.

30. The immunogenic composition of any one of embodiments 22-24 wherein the Gram positive bacteria AI polypeptide is capable of associating with an epithelial cell surface.

31. The immunogenic composition of embodiment 30 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.

32. The immunogenic composition of any of embodiments 22-24 wherein the Gram positive bacteria AI polypeptide is a full-length Gram positive bacteria AI protein.

33. The immunogenic composition of any of embodiments 22-24 wherein the Gram positive bacteria AI polypeptide is a fragment of a full-length Gram positive bacteria AI protein.

34. The immunogenic composition of embodiment 33 wherein the fragment comprises at least 7 contiguous amino acid residues of the Gram positive bacteria AI protein.

35. The immunogenic composition of embodiment 24 wherein the genus *Streptococcus* bacteria is Group A Streptococcus (GAS) bacteria and the Gram positive bacteria AI polypeptide is a GAS AI polypeptide.

36. The immunogenic composition of embodiment 35 wherein the GAS AI polypeptide is selected from a GAS AI-1.

37. The immunogenic composition of embodiment 35 wherein the GAS AI polypeptide is selected from a GAS AI-2.

38. The immunogenic composition of embodiment 35 wherein the GAS AI polypeptide is selected from a GAS AI-3.

39. The immunogenic composition of embodiment 35 wherein the GAS AI polypeptide is selected from a GAS AI-4.

40. The immunogenic composition of any of embodiments 35-39 wherein the GAS AI polypeptide comprises a sortase substrate motif.

41. The immunogenic composition of embodiment 40 wherein the sortase substrate motif is an LPXTG motif.

42. The immunogenic composition of embodiment 41 wherein the LPXTG motif is represented by XXXXG, wherein the X at the first amino acid position is an L, a V, an E, or a Q, wherein the X at the second amino acid position is P if the X at the first amino acid position is an L, the X at the second amino acid position is a V if the X at the first amino acid position is an E or a Q, or the X at the second amino acid position is a V or a P if the X at the first amino acid position is a V, wherein the X at the third amino acid position is any amino acid residue, and wherein the X at the fourth amino acid position is a T if the X at the first amino acid position is a V, an E, or a Q, or the X at the fourth amino acid position is a T, an S, or an A if the X at the first amino acid position is an L.

43. The immunogenic composition of any one of embodiments 35-39 wherein the GAS AI polypeptide affects the ability of GAS bacteria to adhere to epithelial cells.

44. The immunogenic composition of any one of embodiments 35-39 wherein the GAS AI polypeptide affects the ability of GAS bacteria to invade epithelial cells.

45. The immunogenic composition of any one of embodiments 35-39 wherein the GAS AI polypeptide affects the ability of GAS bacteria to translocate through an epithelial cell layer.

46. The immunogenic composition of any one of embodiments 35-39 wherein the GAS AI polypeptide is capable of associating with an epithelial cell surface.



47. The immunogenic composition of embodiment 46 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.

48. The immunogenic composition of any of embodiments 35-39 wherein the GAS AI polypeptide is a full-length GAS AI protein.

49. The immunogenic composition of any of embodiments 35-39 wherein the GAS AI polypeptide is a fragment of a full-length GAS AI protein.

50. The immunogenic composition of embodiment 49 wherein the fragment comprises at least 7 contiguous amino acid residues of the GAS AI protein.

51. The immunogenic composition of embodiment 36 wherein the GAS AI-1 polypeptide is selected from the group consisting of M6\_Spy0157, M6\_Spy0159, M6\_Spy0160, CDC SS 410\_fimbrial, ISS3650\_fimbrial, DSM2071\_fimbrial, and fragments thereof.

52. The immunogenic composition of embodiment 37 wherein the GAS AI-2 polypeptide is selected from the group consisting of GAS15, GAS16, GAS18, and fragments thereof.

53. The immunogenic composition of embodiment 38 wherein the GAS AI-3 polypeptide is selected from the group consisting of SpyM3\_0098, SpyM3\_0100, SpyM3\_0102, SpyM3\_0104, SPs0100, SPs0102, SPs0104, SPs0106, orf78, orf80, orf82, orf84, spyM18\_0126, spyM18\_0128, spyM18\_0130, spyM18\_0132, SpyoM01000156, SpyoM01000155, SpyoM01000154, SpyoM01000153, SpyoM01000152, SpyoM01000151, SpyoM01000150, SpyoM01000149, ISS3040\_fimbrial, ISS3776\_fimbrial, ISS4959\_fimbrial, and fragments thereof.

53. The immunogenic composition of embodiment 39 wherein the GAS AI-4 polypeptide is selected from the group consisting of 19224134, 19224135, 19224137, 19224139, 19224141, 20010296\_fimbrial, 20020069\_fimbrial, CDC SS 635\_fimbrial, ISS4883\_fimbrial, ISS4538\_fimbrial, and fragments thereof.

54. The immunogenic composition of embodiment 24 wherein the *Streptococcus* bacteria is *Streptococcus pneumoniae* and the Gram positive bacteria AI polypeptide is a *S. pneumoniae* AI polypeptide.

55. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide comprises a sortase substrate motif.

56. The immunogenic composition of embodiment 55 wherein the sortase substrate motif is an LPXTG motif.

57. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide affects the ability of *S. pneumoniae* to adhere to epithelial cells.

58. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide affects the ability of *S. pneumoniae* to invade epithelial cells.

59. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide affects the ability of *S. pneumoniae* to translocate through an epithelial cell layer.

60. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide is capable of associating with an epithelial cell surface.

61. The immunogenic composition of embodiment 60 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.

62. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide is a full-length *S. pneumoniae* AI protein.

63. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide is a fragment of a full-length *S. pneumoniae* AI protein.

64. The immunogenic composition of embodiment 63 wherein the fragment comprises at least 7 contiguous amino acid residues of the *S. pneumoniae* AI protein.

65. The immunogenic composition of embodiment 54 wherein the *S. pneumoniae* AI polypeptide is selected from the group consisting of SP0462, SP0463, SP0464, orf3\_670, orf4\_670, orf5\_670, ORF3\_14CSR, ORF4\_14CSR, ORF5\_14CSR, ORF3\_19AH, ORF4\_19AH, ORF5\_19AH, ORF3\_19FTW, ORF4\_19FTW, ORF5\_19FTW, ORF3\_23FP, ORF4\_23FP, ORF5\_23FP, ORF3\_23FTW, ORF4\_23FTW, ORF5\_23FTW, ORF3\_6BF, ORF4\_6BF, ORF5\_6BF, ORF3\_6BSP, ORF4\_6BSP, ORF5\_6BSP, ORF3\_9VSP, ORF4\_9VSP, ORF5\_9VSP, and fragments thereof.

66. The immunogenic composition of any one of embodiments 22-24, 35-39, 51-54, or 65 wherein the oligomeric form is a hyperoligomer.

67. The immunogenic composition of any one of embodiments 22-24, 35-39, 51-54, or 65 further comprising a Gram positive bacteria antigen not associated with an AI.

68. The immunogenic composition of embodiment 67 wherein the antigen is selected from the group consisting of GBS 322 and GBS 276.

69. An immunogenic composition comprising a first and a second Group B Streptococcus (GBS) adhesin island (AI) polypeptide.

70. The immunogenic composition of embodiment 69 wherein a full-length polynucleotide sequence encoding for the first GBS AI polypeptide is not present in a GBS bacteria genome comprising a polynucleotide sequence encoding for the second GBS AI polypeptide.

71. The immunogenic composition of embodiment 69 wherein polynucleotides encoding the first and the second GBS AI polypeptide are each present in genomes of more than one GBS serotype and strain isolate.

72. The immunogenic composition of embodiment 69 wherein the first GBS AI polypeptide is encoded by a GBS AI-1.

73. The immunogenic composition of embodiment 69 wherein the first GBS AI polypeptide is encoded by a GBS AI-2.

74. The immunogenic composition of embodiment 72 wherein the second GBS AI polypeptide is encoded by a GBS AI-2.

75. The immunogenic composition of embodiment 73 wherein the second GBS AI polypeptide is encoded by a GBS AI-2.

76. The immunogenic composition of embodiment 72 wherein the second GBS AI polypeptide is encoded by a GBS AI-1.



77. The immunogenic composition of embodiment 73 wherein the second GBS AI polypeptide is encoded by a GBS AI-1.

78. The immunogenic composition of embodiment 72 wherein the first GBS AI polypeptide is selected from the group consisting of GBS 80, GBS 104, GBS 52, and fragments thereof.

79. The immunogenic composition of embodiment 73 wherein the first GBS AI polypeptide is selected from the group consisting of GBS 59, GBS 67, GBS 150, 01521, 01523, 01524, and fragments thereof.

80. The immunogenic composition of embodiment 74 or 75 wherein the second GBS AI polypeptide is selected from the group consisting of GBS 59, GBS 67, GBS 150, 01521, 01523, 01524, and fragments thereof, and wherein the first and the second GBS AI polypeptide are not the same polypeptide.

81. The immunogenic composition of embodiment 76 or 77 wherein the second GBS AI polypeptide is selected from the group consisting of GBS 80, GBS 104, GBS 52, and fragments thereof, and wherein the first and the second GBS AI polypeptide are not the same polypeptide.

82. The immunogenic composition of any one of embodiments 69-77 wherein the first GBS AI polypeptide comprises a sortase substrate motif.

83. The immunogenic composition of embodiment 82 wherein the sortase substrate motif is an LPXTG motif.

84. The immunogenic composition of embodiment 83 wherein the LPXTG motif is represented by the sequence XPXTG, wherein the X at amino acid position 1 is an L, an I, or an F and the X at amino acid position 3 is any amino acid residue.

85. The immunogenic composition of any one of embodiments 69-77 wherein the first GBS AI polypeptide affects the ability of GBS bacteria to adhere to epithelial cells.

86. The immunogenic composition of any one of embodiments 69-77 wherein the first GBS AI polypeptide affects the ability of GBS bacteria to invade epithelial cells.

87. The immunogenic composition of any one of embodiments 69-77 wherein the first GBS AI polypeptide affects the ability of GBS bacteria to translocate through an epithelial cell layer.

88. The immunogenic composition of any one of embodiments 69-77 wherein the first GBS AI polypeptide is capable of associating with an epithelial cell surface.

89. The immunogenic composition of embodiment 88 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.

90. The immunogenic composition of any of embodiments 69-77 wherein the first GBS AI polypeptide is a full-length GBS AI protein.

91. The immunogenic composition of any of embodiments 69-77 wherein the first GBS AI polypeptide is a fragment of a full-length GBS AI protein.

92. The immunogenic composition of embodiment 91 wherein the fragment comprises at least 7 contiguous amino acid residues of the first GBS AI protein.

93. The immunogenic composition of any one of embodiments 69-79 wherein the first GBS AI polypeptide is in oligomeric form.

94. The immunogenic composition of any one of embodiments 69-77 wherein the second GBS AI polypeptide is in oligomeric form.

95. The immunogenic composition of any one of embodiments 69-79 wherein the first and the second GBS AI polypeptide are associated in a single oligomeric form.

96. The immunogenic composition of embodiment 95 wherein the first and the second GBS AI polypeptides are chemically associated.

97. The immunogenic composition of embodiment 95 wherein the first and the second GBS AI polypeptides are physically associated.

98. The immunogenic composition of embodiment 93 wherein the oligomeric form is a hyperoligomer.

99. The immunogenic composition of embodiment 94 wherein the oligomeric form is a hyperoligomer.

100. The immunogenic composition of embodiment 76 wherein the first GBS AI polypeptide is GBS 80 and the second GBS AI polypeptide is GBS 104.

101. The immunogenic composition of embodiment 74 wherein the first GBS AI polypeptide is GBS 80 and the second GBS AI polypeptide is GBS 67.

102. The immunogenic composition of any one of embodiments 69-79, 100, or 101 further comprising a GBS polypeptide not associated with an AI.

103. The immunogenic composition of embodiment 102 wherein the GBS polypeptide not associated with an AI is selected from the group consisting of GBS 322 and GBS 276.

104. The immunogenic composition of embodiment 103 wherein the GBS polypeptide not associated with an AI is GBS 322.

105. An immunogenic composition comprising a first and a second Gram positive bacteria adhesin island (AI) polypeptide.

106. The immunogenic composition of embodiment 105 wherein a full length polynucleotide sequence encoding for the first Gram positive bacteria AI polypeptide is not present in a genome of a Gram positive bacteria comprising a full length polynucleotide sequence encoding for the second Gram positive bacteria AI polypeptide.

107. The immunogenic composition of embodiment 105 wherein polynucleotides encoding the first and the second Gram positive bacteria AI polypeptide are each present in genomes of more than one Gram positive bacteria serotype and strain isolate.

108. The immunogenic composition of embodiment 105 wherein the first and the second Gram positive bacteria AI polypeptides are of different Gram positive bacteria species.

109. The immunogenic composition of embodiment 105 wherein the first and the second Gram positive bacteria AI polypeptides are of the same Gram positive bacteria species.



10. The immunogenic composition of embodiment 105 wherein the first and the second Gram positive bacteria AI polypeptides are from different AI subtypes.

111. The immunogenic composition of embodiment 105 wherein the first and the second Gram positive bacteria AI polypeptides are from the same AI subtype.

5 112. The immunogenic composition of embodiment 105 wherein the first Gram positive bacteria AI polypeptide has detectable surface exposure on a first Gram positive bacteria strain or serotype but not a second Gram positive bacteria strain or subtype and the second Gram positive bacteria AI polypeptide has detectable surface exposure on the second Gram positive bacteria strain or serotype but not the first Gram positive bacteria strain or serotype.

10 113. The immunogenic composition of embodiment 105 wherein the Gram positive bacteria is *S. pneumoniae*, *S. mutans*, *E. faecalis*, *E. faecium*, *C. difficile*, *L. monocytogenes*, or *C. diphtheriae*.

114. The immunogenic composition of any of embodiments 105-113 wherein the first and the second Gram positive bacteria AI polypeptides comprise a sortase substrate motif.

15 115. The immunogenic composition of embodiment 114 wherein the sortase substrate motif is an LPXTG motif.

20 116. The immunogenic composition of embodiment 115 wherein the LPXTG motif is represented by XXXXG, wherein the X at amino acid position 1 is an L, a V, an E, an I, an F, or a Q, wherein X at amino acid position 2 is a P if X at amino acid position 1 is an L, an I, or an F, wherein X at amino acid position 2 is a V if X at amino acid position 1 is a E or a Q, wherein X at amino acid position 2 is a V or a P if X at amino acid position 1 is a V, wherein X at amino acid position 3 is any amino acid residue, wherein X at amino acid position 4 is a T if X at amino acid position 1 is a V, E, I, F, or Q, and wherein X at amino acid position 4 is a T, S, or A if X at amino acid position 1 is an L.

117. The immunogenic composition of embodiment 105 wherein the first Gram positive bacteria AI polypeptide is a first Group A Streptococcus (GAS) AI polypeptide.

25 118. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide comprises a sortase substrate motif.

119. The immunogenic composition of embodiment 118 wherein the sortase substrate motif is an LPXTG motif.

30 120. The immunogenic composition of embodiment 119 wherein the LPXTG motif is represented by XXXXG, wherein the X at the first amino acid position is an L, a V, an E, or a Q, wherein the X at the second amino acid position is P if the X at the first amino acid position is an L, the X at the second amino acid position is a V if the X at the first amino acid position is an E or a Q, or the X at the second amino acid position is a V or a P if the X at the first amino acid position is a V, wherein the X at the third amino acid position is any amino acid residue, and wherein the X at the fourth amino acid position is a T if the X at the first amino acid position is a V, an E, or a Q, or the X at the fourth amino acid position is a T, an S, or an A if the X at the first amino acid position is an L.

35 121. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide affects the ability of GAS bacteria to adhere to epithelial cells.

122. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide affects the ability of GAS bacteria to invade epithelial cells.

123. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide affects the ability of GAS bacteria to translocate through an epithelial cell layer.

124. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide is capable of associating with an epithelial cell surface.

125. The immunogenic composition of embodiment 117 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.

126. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide is a full-length GAS AI protein.

127. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide is a fragment of a full-length GAS AI protein.

128. The immunogenic composition of embodiment 127 wherein the fragment comprises at least 7 contiguous amino acid residues of the GAS AI protein.

129. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide is a first GAS AI-1 polypeptide.

130. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide is a first GAS AI-2 polypeptide.

131. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide is a first GAS AI-3 polypeptide.

132. The immunogenic composition of embodiment 117 wherein the first GAS AI polypeptide is a first GAS AI-4 polypeptide.

133. The immunogenic composition of any one of embodiments 117 or 129-132 wherein the second Gram positive bacteria AI polypeptide is a second GAS AI polypeptide.

134. The immunogenic composition of embodiment 133 wherein the second GAS AI polypeptide is a second GAS AI-1 polypeptide.

135. The immunogenic composition of embodiment 133 wherein the second GAS AI polypeptide is a second GAS AI-2 polypeptide.

136. The immunogenic composition of embodiment 133 wherein the second GAS AI polypeptide is a second GAS AI-3 polypeptide.

137. The immunogenic composition of embodiment 133 wherein the second GAS AI polypeptide is a second GAS AI-4 polypeptide.

138. The immunogenic composition of embodiment 129 wherein the first GAS AI-1 polypeptide is selected from the group consisting of M6\_Spy0157, M6\_Spy0159, M6\_Spy0160, CDC SS 410\_fimbrial, ISS3650\_fimbrial, DSM2071\_fimbrial, and fragments thereof.

139. The immunogenic composition of embodiment 130 wherein the first GAS AI-2 polypeptide is selected from the group consisting of GAS15, GAS16, GAS18, and fragments thereof.



140. The immunogenic composition of embodiment 131 wherein the first GAS AI-3 polypeptide is selected from the group consisting of SpyM3\_0098, SpyM3\_0100, SpyM3\_0102, SpyM3\_0104, SPs0100, SPs0102, SPs0104, SPs0106, orf78, orf80, orf82, orf84, spyM18\_0126, spyM18\_0128, spyM18\_0130, spyM18\_0132, SpyoM01000156, SpyoM01000155, SpyoM01000154, SpyoM01000153, SpyoM01000152, SpyoM01000151, SpyoM01000150, SpyoM01000149, ISS3040\_fimbrial, ISS3776\_fimbrial, ISS4959\_fimbrial, and fragments thereof.

141. The immunogenic composition of embodiment 132 wherein the first GAS AI-4 polypeptide is selected from the group consisting of 19224134, 19224135, 19224137, 19224139, 19224141, 20010296\_fimbrial, 20020069\_fimbrial, CDC SS 635\_fimbrial, ISS4883\_fimbrial, ISS4538\_fimbrial, and fragments thereof.

142. The immunogenic composition of embodiment 134 wherein the second GAS AI-1 polypeptide is selected from the group consisting of M6\_Spy0157, M6\_Spy0159, M6\_Spy0160, CDC SS 410\_fimbrial, ISS3650\_fimbrial, DSM2071\_fimbrial, and fragments thereof.

143. The immunogenic composition of embodiment 135 wherein the second GAS AI-2 polypeptide is selected from the group consisting of GAS15, GAS16, GAS18, and fragments thereof.

144. The immunogenic composition of embodiment 136 wherein the second GAS AI-3 polypeptide is selected from the group consisting of SpyM3\_0098, SpyM3\_0100, SpyM3\_0102, SpyM3\_0104, SPs0100, SPs0102, SPs0104, SPs0106, orf78, orf80, orf82, orf84, spyM18\_0126, spyM18\_0128, spyM18\_0130, spyM18\_0132, SpyoM01000156, SpyoM01000155, SpyoM01000154, SpyoM01000153, SpyoM01000152, SpyoM01000151, SpyoM01000150, SpyoM01000149, ISS3040\_fimbrial, ISS3776\_fimbrial, ISS4959\_fimbrial, and fragments thereof.

145. The immunogenic composition of embodiment 137 wherein the second GAS AI-4 polypeptide is selected from the group consisting of 19224134, 19224135, 19224137, 19224139, 19224141, 20010296\_fimbrial, 20020069\_fimbrial, CDC SS 635\_fimbrial, ISS4883\_fimbrial, ISS4538\_fimbrial, and fragments thereof.

146. The immunogenic composition of any one of embodiments 117-132 or 138-141 wherein the second Gram positive bacteria AI polypeptide is a Group B Streptococcus (GBS) AI polypeptide.

147. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide comprises a sortase substrate motif.

148. The immunogenic composition of embodiment 147 wherein the sortase substrate motif is an LPXTG motif.

149. The immunogenic composition of embodiment 148 wherein the LPXTG motif is represented by the amino acid sequence XPXTG, wherein the X at amino acid position 1 is an L, an I, or an F and the X at amino acid position 3 is any amino acid residue.

150. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide affects the ability of GBS bacteria to adhere to epithelial cells.

151. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide affects the ability of GBS bacteria to invade epithelial cells.

152. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide affects the ability of GBS bacteria to translocate through an epithelial cell layer.

153. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide is capable of associating with an epithelial cell surface.

5 154. The immunogenic composition of embodiment 146 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.

155. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide is a full-length GBS AI protein.

10 156. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide is a fragment of a full-length GBS AI protein.

157. The immunogenic composition of embodiment 156 wherein the fragment comprises at least 7 contiguous amino acid residues of the GBS AI protein.

158. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide is a GBS AI-1 polypeptide.

15 159. The immunogenic composition of embodiment 146 wherein the GBS AI polypeptide is a GBS AI-2 polypeptide.

160. The immunogenic composition of embodiment 158 wherein the GBS AI-1 polypeptide is selected from the group consisting of GBS 80, GBS 104, GBS 52, and fragments thereof.

20 161. The immunogenic composition of embodiment 159 wherein the GBS AI-2 polypeptide is selected from the group consisting of GBS 59, GBS 67, GBS 150, 01521, 01523, 01524, and fragments thereof.

162. The immunogenic composition of any one of embodiments 117-132 or 138-141 wherein the second Gram positive bacteria AI polypeptide is a *Streptococcus pneumoniae* AI polypeptide.

25 163. The immunogenic composition of embodiment 162 wherein the *S. pneumoniae* AI polypeptide comprises a sortase substrate motif.

164. The immunogenic composition of embodiment 163 wherein the sortase substrate motif is an LPXTG motif.

165. The immunogenic composition of embodiment 162 wherein the *S. pneumoniae* AI polypeptide affects the ability of *S. pneumoniae* to adhere to epithelial cells.

30 166. The immunogenic composition of embodiment 162 *S. pneumoniae* AI polypeptide affects the ability of *S. pneumoniae* to invade epithelial cells.

167. The immunogenic composition of embodiment 162 wherein the *S. pneumoniae* AI polypeptide affects the ability of *S. pneumoniae* to translocate through an epithelial cell layer.

35 168. The immunogenic composition of embodiment 162 wherein the *S. pneumoniae* AI polypeptide is capable of associating with an epithelial cell surface.

169. The immunogenic composition of embodiment 168 wherein the associating with an epithelial cell surface is binding to the epithelial cell surface.



170. The immunogenic composition of embodiment 162 wherein the *S. pneumoniae* AI polypeptide is a full-length *S. pneumoniae* AI protein.

171. The immunogenic composition of embodiment 162 wherein the *S. pneumoniae* AI polypeptide is a fragment of a full-length *S. pneumoniae* AI protein.

172. The immunogenic composition of embodiment 162 wherein the fragment comprises at least 7 contiguous amino acid residues of the *S. pneumoniae* AI protein.

173. The immunogenic composition of embodiment 162 wherein the *S. pneumoniae* AI polypeptide is selected from the group consisting of SP0462, SP0463, SP0464, orf3\_670, orf4\_670, orf5\_670, ORF3\_14CSR, ORF4\_14CSR, ORF5\_14CSR, ORF3\_19AH, ORF4\_19AH, ORF5\_19AH, ORF3\_19FTW, ORF4\_19FTW, ORF5\_19FTW, ORF3\_23FP, ORF4\_23FP, ORF5\_23FP, ORF3\_23FTW, ORF4\_23FTW, ORF5\_23FTW, ORF3\_6BF, ORF4\_6BF, ORF5\_6BF, ORF3\_6BSP, ORF4\_6BSP, ORF5\_6BSP, ORF3\_9VSP, ORF4\_9VSP, ORF5\_9VSP, and fragments thereof.

174. The immunogenic composition of any one of embodiments 105-117 wherein the first Gram positive bacteria AI polypeptide is in oligomeric form.

175. The immunogenic composition of embodiment 174 wherein the oligomeric form is a hyperoligomer.

176. The immunogenic composition of embodiment 174 wherein the second Gram positive bacteria AI polypeptide is in oligomeric form.

177. The immunogenic composition of embodiment 176 wherein the oligomeric form is a hyperoligomer.

178. The immunogenic composition of embodiment 176 wherein the first and the second Gram positive bacteria AI polypeptide are associated in a single oligomeric form.

179. The immunogenic composition of embodiment 178 wherein the first and the second Gram positive bacteria AI polypeptide are chemically associated.

180. The immunogenic composition of embodiment 178 wherein the first and the second Gram positive bacteria AI polypeptide are physically associated.

181. The immunogenic composition of any one of embodiments 105-117 further comprising a Gram positive bacteria polypeptide not associated with an AI.

182. The immunogenic composition of embodiment 181 wherein the Gram positive bacteria polypeptide not associated with an AI is selected from the group consisting of GBS 322 and GBS 276.

183. The immunogenic composition of embodiment 182 wherein the Gram positive bacteria polypeptide not associated with an AI is GBS 322.

184. A modified Gram positive bacterium adapted to produce increased levels of AI surface protein.

185. The modified Gram positive bacterium of embodiment 184 wherein the AI surface protein is in oligomeric form.

186. The modified Gram positive bacterium of embodiment 185 wherein the oligomeric form is a hyperoligomer.

187. The modified Gram positive bacterium of any one of embodiments 184-186 which is a Group B Streptococcus bacterium.

188. The modified Gram positive bacterium of any one of embodiments 184-186 which is a Group A Streptococcus bacterium.

5 189. The modified Gram positive bacterium of any one of embodiments 184-186 which is a non-pathogenic Gram positive bacterium.

190. The modified Gram positive bacterium of embodiment 189 wherein the non-pathogenic Gram positive bacterium is *Streptococcus gordonii*.

10 191. The modified Gram positive bacterium of embodiment 189 wherein the non-pathogenic Gram positive bacterium is *Lactococcus lactis*.

192. The modified Gram positive bacterium of any one of embodiments 184-186 which has been inactivated and wherein the AI surface protein is exposed on the surface of the Gram positive bacterium.

15 193. The modified Gram positive bacterium of any one of embodiments 184-186 which has been attenuated and wherein the AI surface protein is exposed on the surface of the Gram positive bacterium.

194. The modified GBS bacterium of embodiment 187 which has been inactivated and wherein the AI surface protein is exposed on the surface of the GBS bacterium.

20 195. The modified GBS bacterium of embodiment 187 which has been attenuated and wherein the AI surface protein is exposed on the surface of the GBS bacterium.

196. The modified GAS bacterium of embodiment 188 which has been inactivated and wherein the AI surface protein is exposed on the surface of the GAS bacterium.

197. The modified GAS bacterium of embodiment 188 which has been attenuated and wherein the AI surface protein is exposed on the surface of the GAS bacterium.

25 198. The modified non-pathogenic bacterium of embodiment 189 which has been inactivated and wherein the AI surface protein is exposed on the surface of the non-pathogenic Gram positive bacterium.

30 199. The modified non-pathogenic bacterium of embodiment 189 which has been attenuated and wherein the AI surface protein is exposed on the surface of the non-pathogenic Gram positive bacterium.

200. A method for manufacturing an oligomeric adhesin island (AI) surface antigen comprising:

culturing a Gram positive bacterium that expresses an oligomeric AI surface antigen and isolating the expressed oligomeric AI surface antigen.

35 201. The method of embodiment 200 wherein the step of isolating is performed by collecting said oligomeric AI surface antigen from Gram positive bacterium secretions in the Gram positive bacterium culture.

202. The method of embodiment 200 further comprising a step of purifying.



203. The method of embodiment 202 wherein the oligomeric AI surface antigen is purified from the Gram positive bacterium cell surface.

204. The method of embodiment 200 wherein the Gram positive bacterium is adapted for increased AI protein expression.

5 205. The method of any one of embodiments 200-204 wherein the Gram positive bacterium is a Group A Streptococcus bacterium.

206. The method of any one of embodiments 200-204 wherein the Gram positive bacterium is a Group B Streptococcus bacterium.

10 207. The method of any one of embodiments 200-204 wherein the oligomeric AI surface antigen is in hyperoligomeric form.

208. The method of embodiment 200 wherein the Gram positive bacterium expresses the oligomeric AI surface antigen recombinantly.

209. The method of embodiment 208 wherein the Gram positive bacterium further manipulated expresses at least 1 AI sortase.

15 210. The modified Gram positive bacterium of any one of embodiments 184-186 which is a *S. pneumoniae* bacterium.

211. The method of any one of embodiments 200-204 wherein the Gram positive bacterium is *S. pneumoniae*.

1. An immunogenic composition comprising a purified Group B Streptococcus (GBS) adhesin island (AI) polypeptide in oligomeric form.

5 2. The immunogenic composition of claim 1 wherein the GBS AI polypeptide is selected from a GBS AI-1.

3. The immunogenic composition of claim 1 wherein the GBS AI polypeptide is selected from a GBS AI-2.

4. The immunogenic composition of claim 2 wherein the GBS AI polypeptide is selected from the group consisting of GBS 80, GBS 104, GBS 52, and fragments thereof.

10 5. The immunogenic composition of claim 3 wherein the GBS AI polypeptide is selected from the group consisting of GBS 59, GBS 67, GBS 150, 01521, 01523, 01524, and fragments thereof.

6. The immunogenic composition of claim 4 wherein the GBS AI polypeptide is GBS 80.

15 7. The immunogenic composition of any of claims 1-6 wherein the oligomeric form is a hyperoligomer.

8 (22). An immunogenic composition comprising a purified Gram positive bacteria adhesin island (AI) polypeptide in an oligomeric form.

20 9 (23). The immunogenic composition of claim 8 wherein the Gram positive bacteria is of a genus selected from the group consisting of *Streptococcus*, *Enterococcus*, *Staphylococcus*, *Clostridium*, *Corynebacterium*, or *Listeria*.

10 (24). The immunogenic composition of claim 9 wherein the Gram positive bacteria is of the genus *Streptococcus*.

25 11 (35). The immunogenic composition of claim 10 wherein the genus *Streptococcus* bacteria is Group A Streptococcus (GAS) bacteria and the Gram positive bacteria AI polypeptide is a GAS AI polypeptide.

12 (36). The immunogenic composition of claim 11 wherein the GAS AI polypeptide is selected from a GAS AI-1.

13 (37). The immunogenic composition of claim 11 wherein the GAS AI polypeptide is selected from a GAS AI-2.

30 14 (38). The immunogenic composition of claim 11 wherein the GAS AI polypeptide is selected from a GAS AI-3.

15 (39). The immunogenic composition of claim 11 wherein the GAS AI polypeptide is selected from a GAS AI-4.

35 16 (66). The immunogenic composition of any one of claims 8-15 wherein the oligomeric form is a hyperoligomer.

17. An immunogenic composition comprising a first and a second Group B Streptococcus (GBS) adhesin island (AI) polypeptide.



18. The immunogenic composition of claim 17 wherein the first GBS AI polypeptide is encoded by a GBS AI-1.

19. The immunogenic composition of claim 18 wherein the second GBS AI polypeptide is encoded by a GBS AI-2.

20. The immunogenic composition of claim 18 wherein the first GBS AI polypeptide is selected from the group consisting of GBS 80, GBS 104, GBS 52, and fragments thereof.

21. The immunogenic composition of claim 19 wherein the second GBS AI polypeptide is selected from the group consisting of GBS 59, GBS 67, GBS 150, 01521, 01523, 01524, and fragments thereof, and wherein the first and the second GBS AI polypeptide are not the same polypeptide.

22. The immunogenic composition of claim 19 wherein the first GBS AI polypeptide is GBS 80 and the second GBS AI polypeptide is GBS 67.

23. An immunogenic composition comprising a first and a second Gram positive bacteria adhesin island (AI) polypeptide.

24. The immunogenic composition of claim 23 wherein the Gram positive bacteria is *Streptococcus*, *Enterococcus*, *Staphylococcus*, *Clostridium*, *Corynebacterium*, or *Listeria*.

25. The immunogenic composition of claim 23 wherein the first Gram positive bacteria AI polypeptide is a first Group A Streptococcus (GAS) AI polypeptide.

26. The immunogenic composition of claim 25 wherein the first GAS AI polypeptide is a first GAS AI-1 polypeptide.

27. The immunogenic composition of claim 25 wherein the first GAS AI polypeptide is a first GAS AI-2 polypeptide.

28. The immunogenic composition of claim 25 wherein the first GAS AI polypeptide is a first GAS AI-3 polypeptide.

29. The immunogenic composition of claim 25 wherein the first GAS AI polypeptide is a first GAS AI-4 polypeptide.

30. The immunogenic composition of any one of claims 25-29 wherein the second Gram positive bacteria AI polypeptide is a second GAS AI polypeptide.

31. The immunogenic composition of claim 30 wherein the second GAS AI polypeptide is a second GAS AI-1 polypeptide.

32. The immunogenic composition of claim 30 wherein the second GAS AI polypeptide is a second GAS AI-2 polypeptide.

33. The immunogenic composition of claim 30 wherein the second GAS AI polypeptide is a second GAS AI-3 polypeptide.

34. The immunogenic composition of claim 30 wherein the second GAS AI polypeptide is a second GAS AI-4 polypeptide.

35. A modified Gram positive bacterium adapted to produce increased levels of AI surface protein.

36. The modified Gram positive bacterium of claim 35 wherein the AI surface protein is in oligomeric form.

37. The modified Gram positive bacterium of claim 36 wherein the oligomeric form is a hyperoligomer.

5 38. The modified Gram positive bacterium of any one of claims 35-37 which is a non-pathogenic Gram positive bacterium.

39. The modified Gram positive bacterium of claim 38 wherein the non-pathogenic Gram positive bacterium is *Lactococcus lactis*.

10 40. A method for manufacturing an oligomeric adhesin island (AI) surface antigen comprising:

culturing a Gram positive bacterium that expresses an oligomeric AI surface antigen and isolating the expressed oligomeric AI surface antigen.



FIGURE 1: Adhesion Island 1

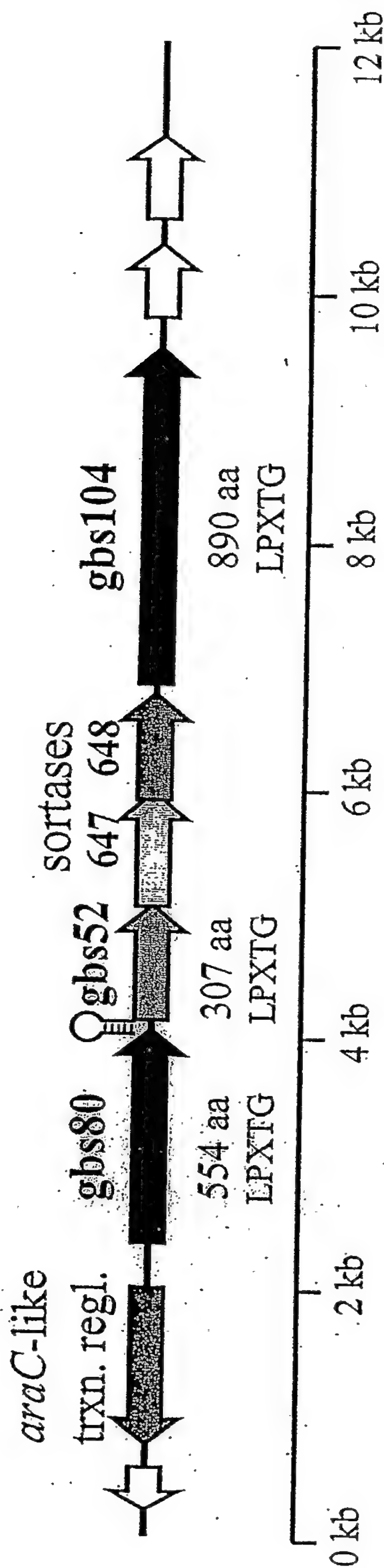
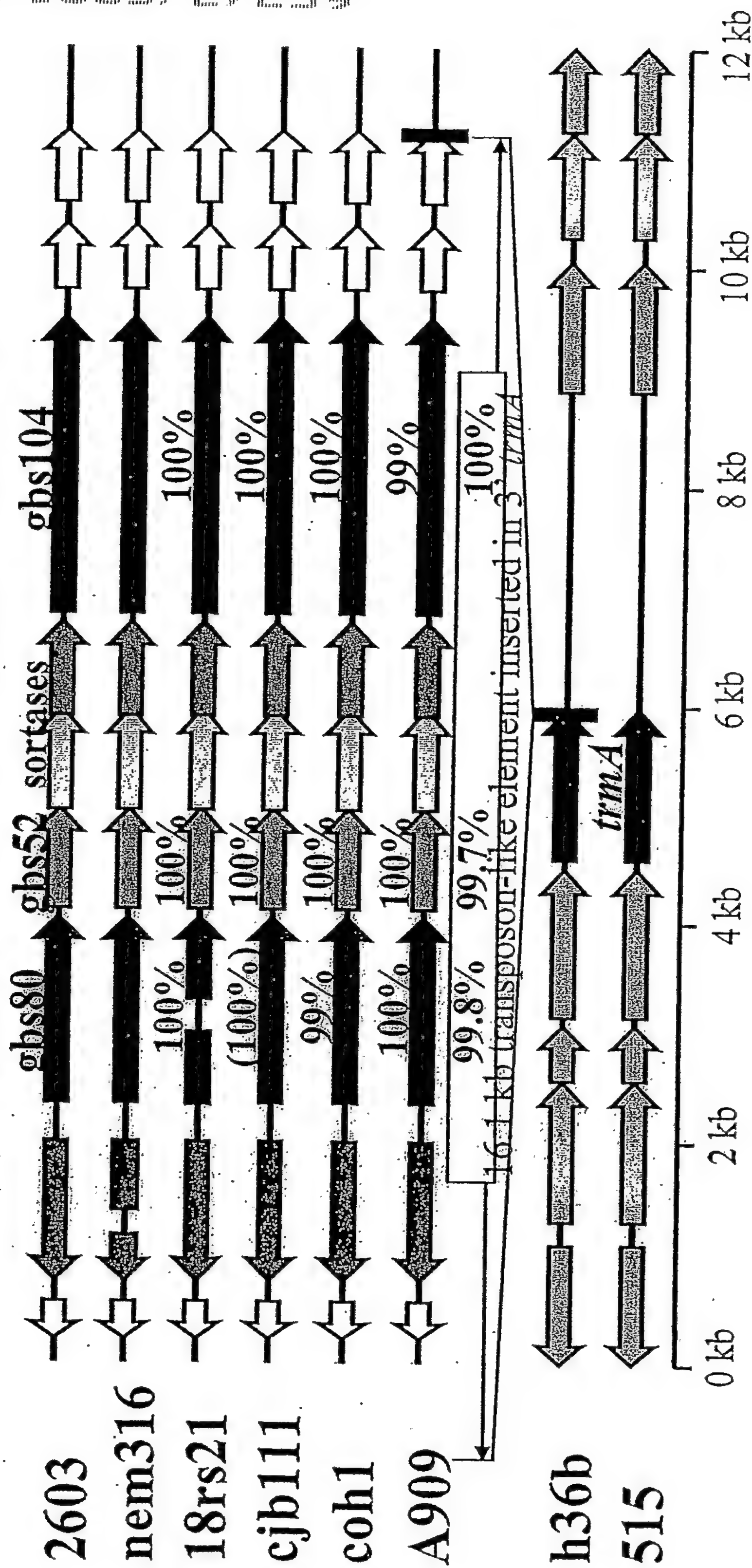


Figure 2: Conservation of AI-1 in GBS serotypes and strain isolates





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FIGURE 3: Correlation of AI-1 and AI-2 within GBS serotype V, strain isolate 2603 genome

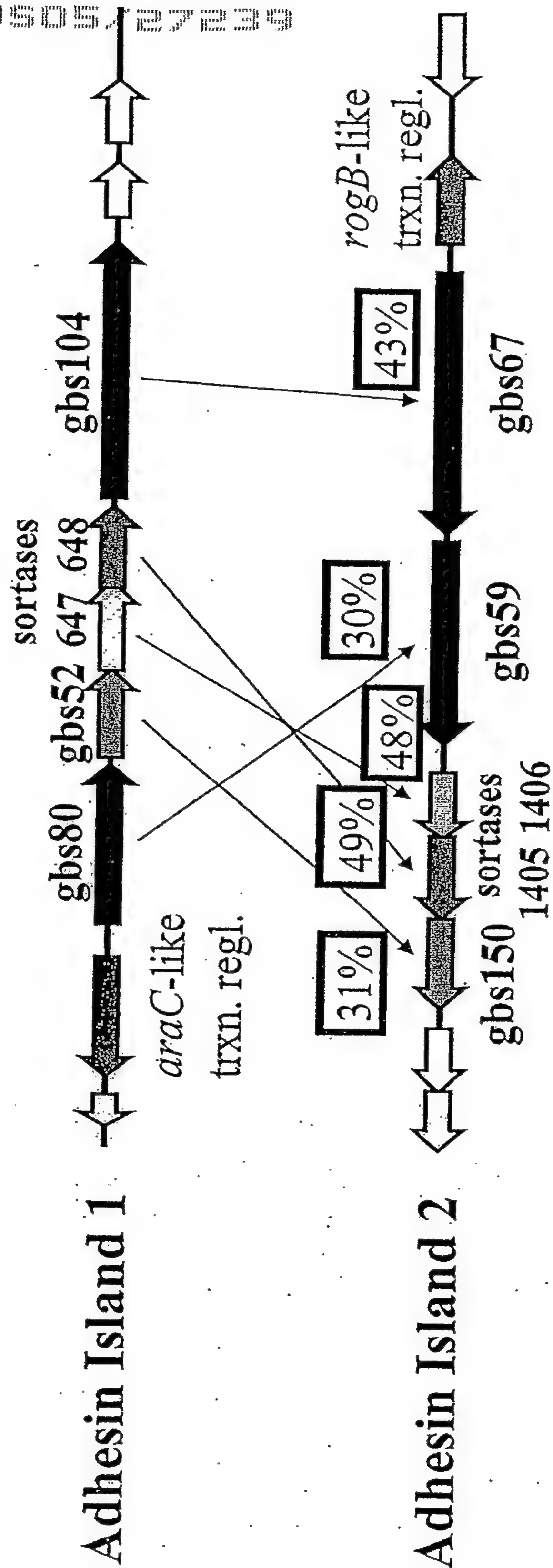
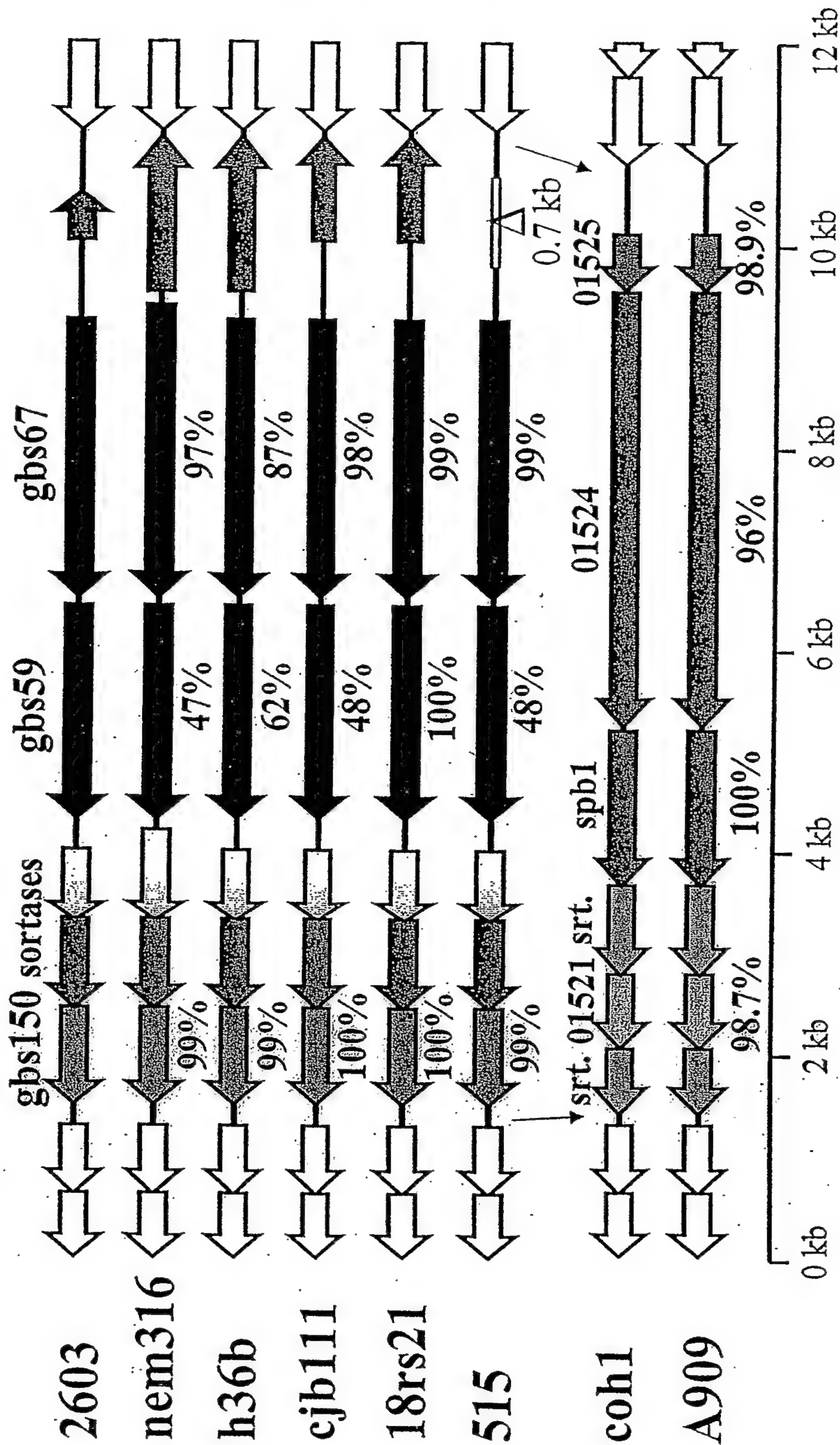
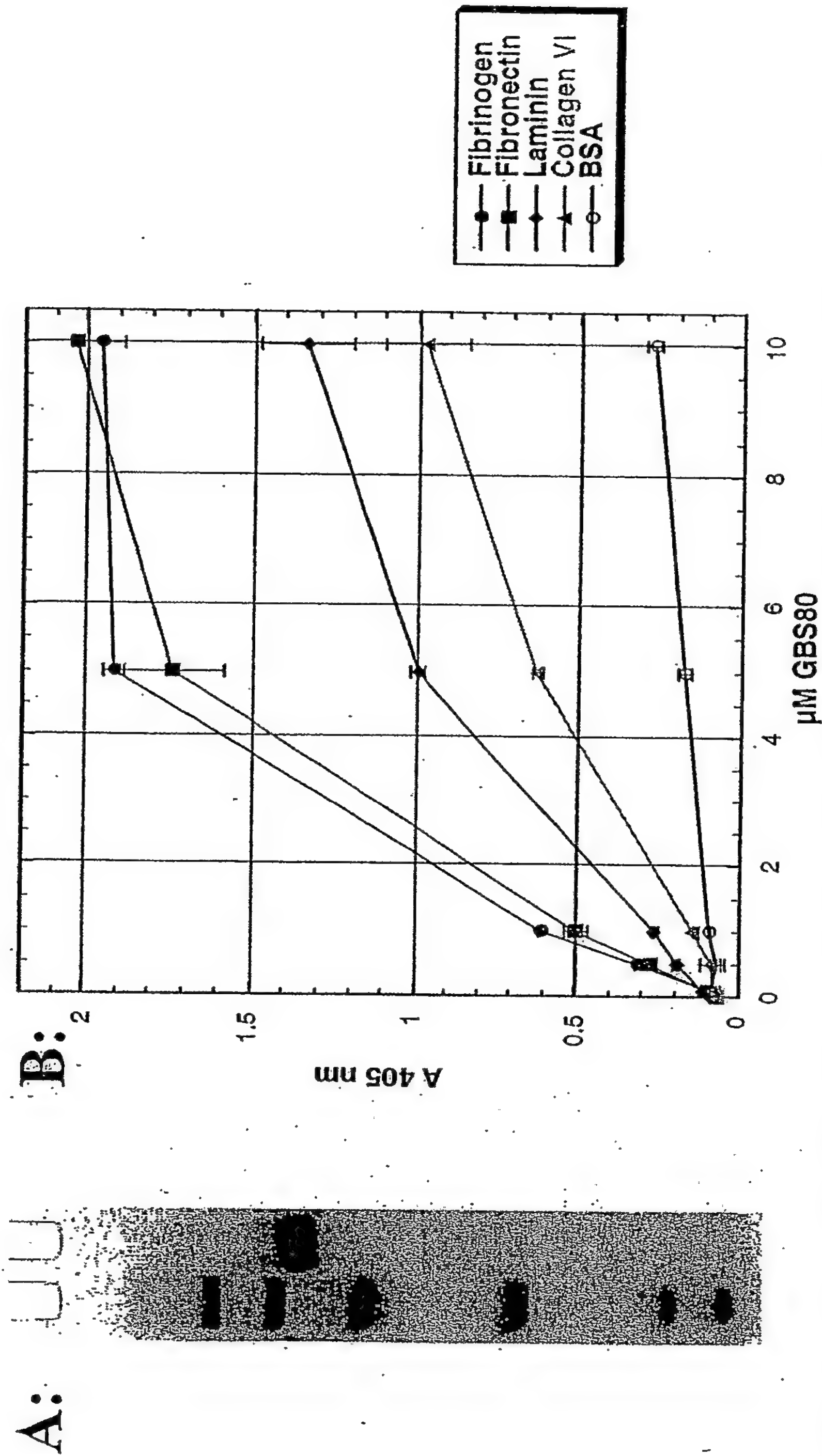


Figure 4: Identification and Variance of AI-2 in Several GBS Serotypes and Strain Isolates



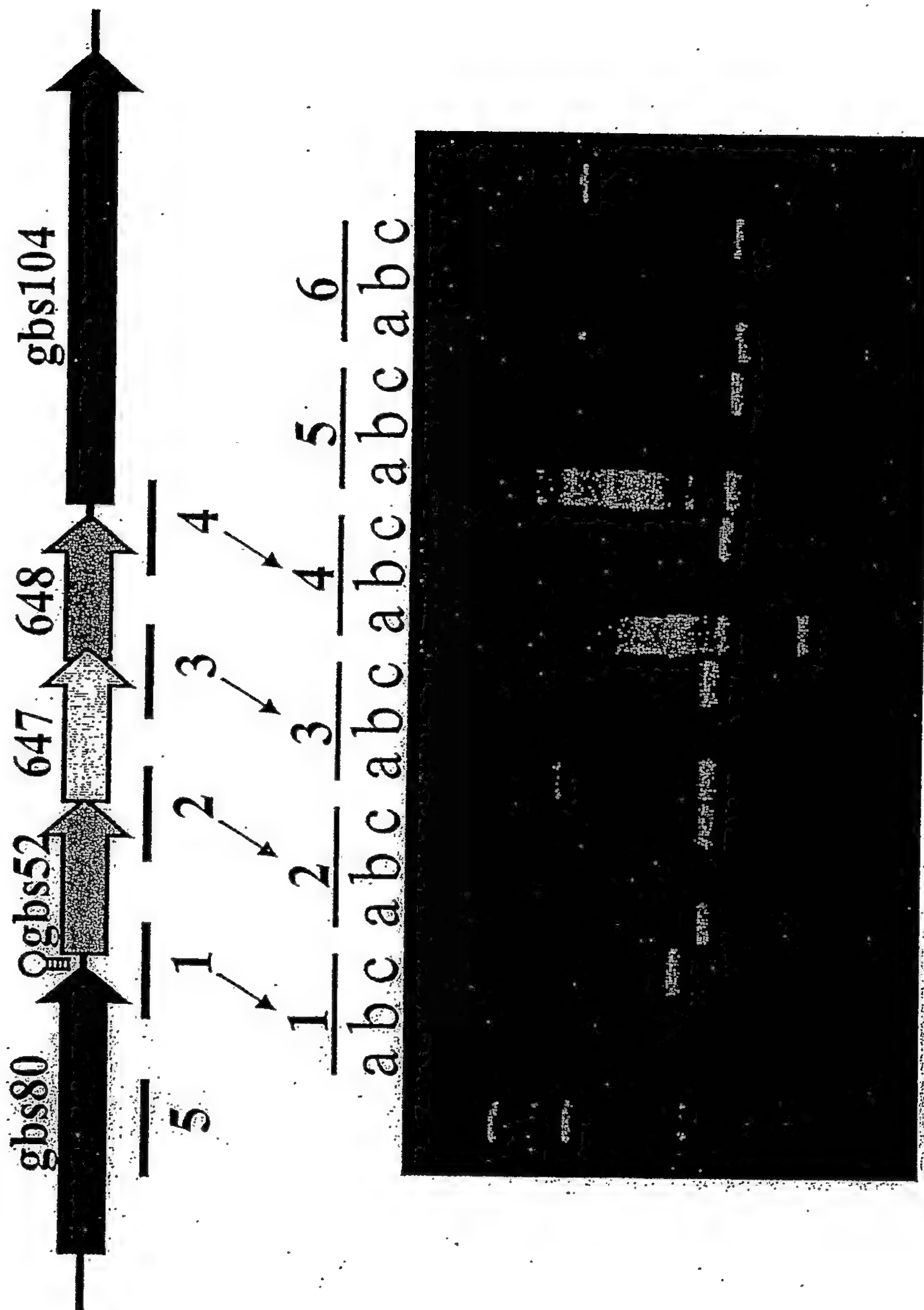


**Figure 5: Purified gbs80 protein binds fibronectin and fibrinogen in an ELISA**

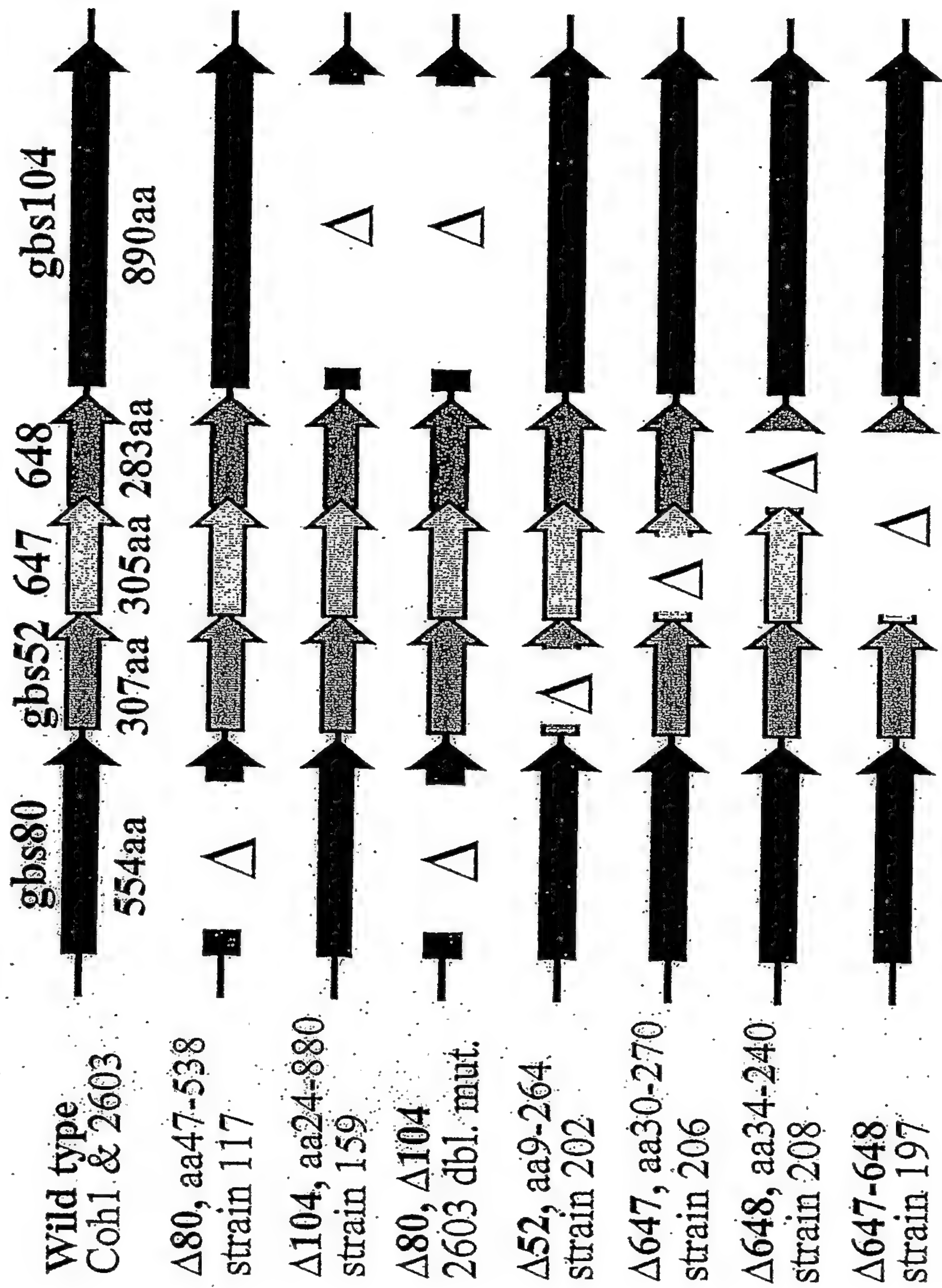


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Figure 6: Adhesion Island I is an operon by RT-PCR





**Figure 7: In frame deletions of AI-1 genes**

**Figure 8: gbs80 is required for surface localization of gbs104**

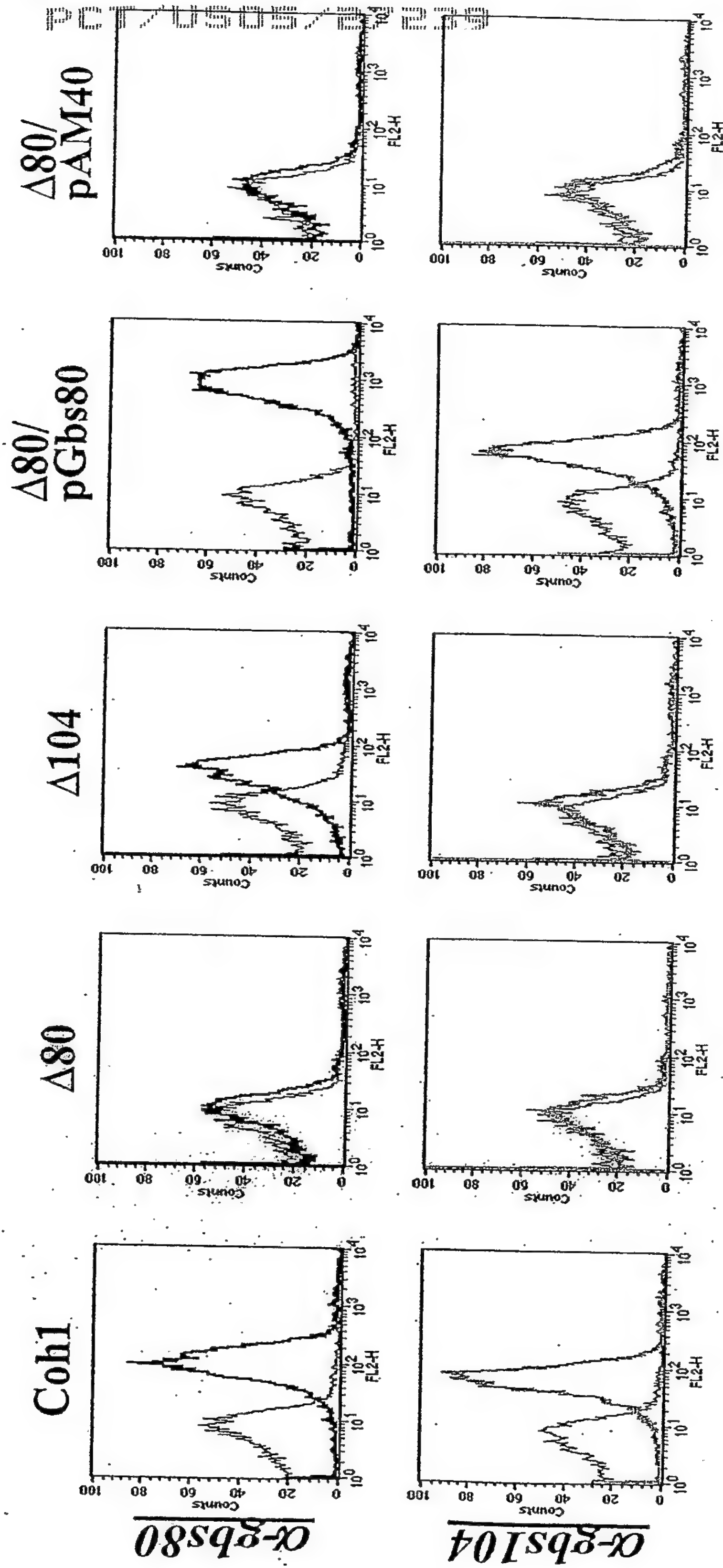
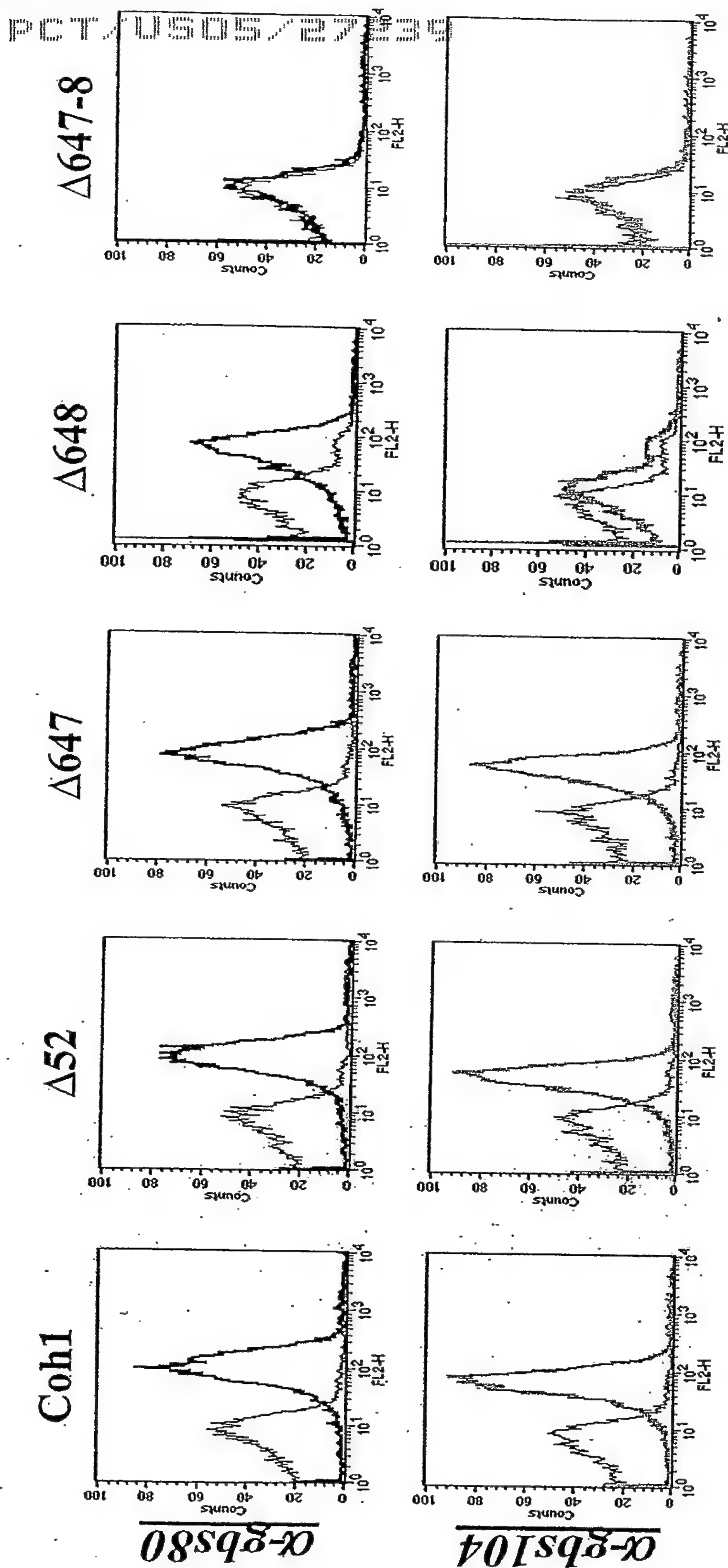
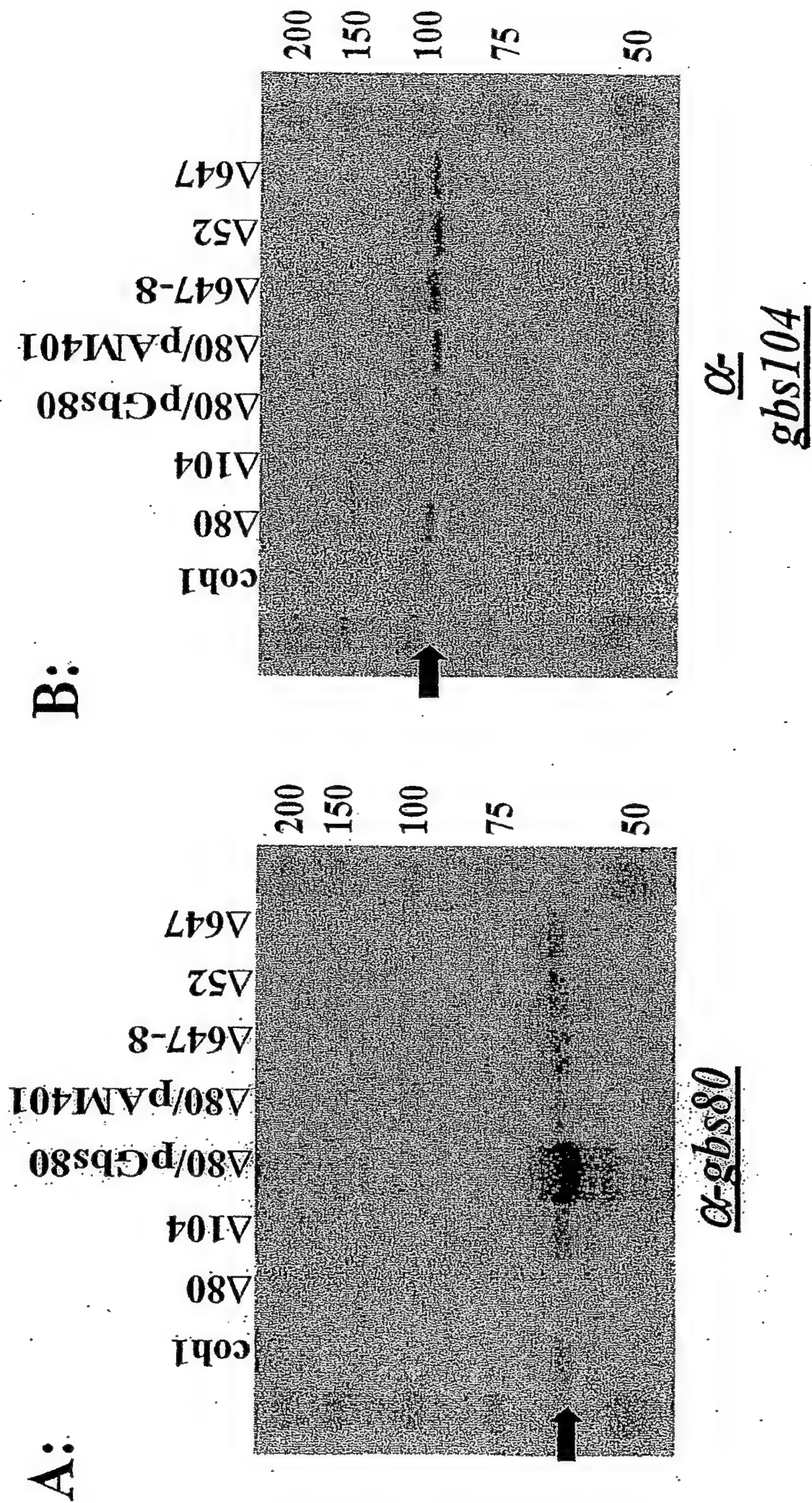


Figure 9: sortases 647 & 648 play a semi-redundant role in surface exposure of gbs80 and gbs104



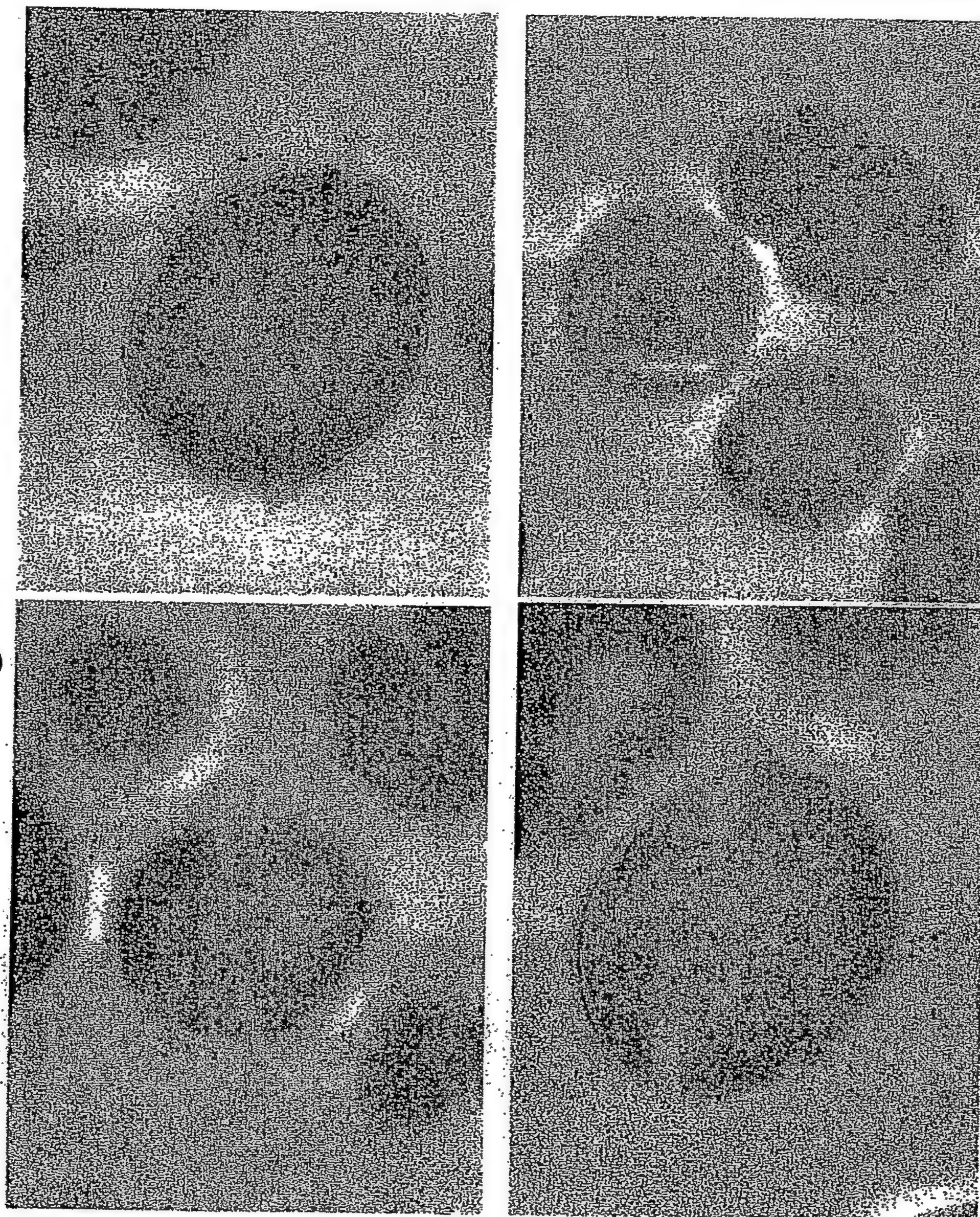


**Figure 10: Western blot of mutant strains**





**FIGURE 11: Pre-embedding IEM  
staining of GBS 80**





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FIGURE 12: Predicted Secondary Structure for GBS 067

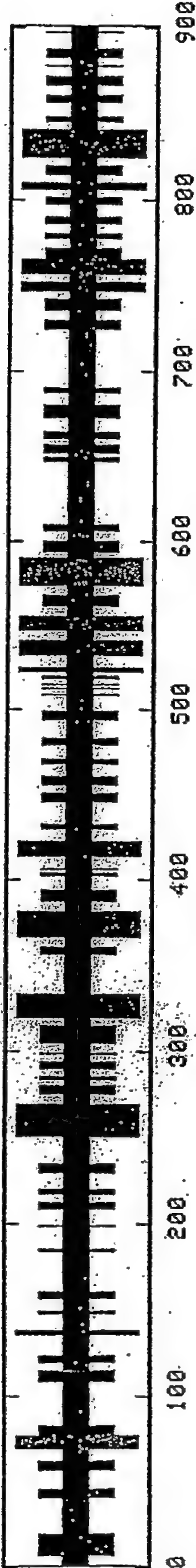
PHD SECONDARY STRUCTURE PREDICTION for GBS 067

10 20 30 40 50 60 70  
MRKYQFSKILTLFLSLQPLNTNVLGSESTVPENGAKGLVVKTTDDQNKPLSKATFVLKTTAHPESK  
CC  
IEKVTAEITGEATFDNLIPGDYTLSEETAPEGYKKTNTQWQVKSNGKTTIQNSGDKNSTIGNQEELD  
hHHhhheeecc  
KQYPTGIYEDTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRISSEVGDLAHNKYKI  
CC  
ELTVSGKTIKPVKQKPLDVVFLDNSNSMNDGPNFQRHNKAKAAEALGTAVKDILGANSNDNRVALV  
EEEECC  
TYGSDIFDGRSVDVVKGFKEDDKYYGLQTKFTIQTENYSHKQLTNNAEEIIKRIPTEAPKAKWGSTTNGL  
EccceecCCeeEeccecccccceeeEeeEeccccchHHHHHHHHHHhhcccccccccccccccc  
TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTGVPTRSYAINNFKLGAYESQFEQM  
CC  
KKNGLYLNKSNFLITDKPEDIKNGESYFLPLDSYQTQIISGNLQKLHYLDNLNYPKGTIYRNGVPVKEH  
HHhCC  
GTPTKLYINSLKQKNYDIFNFGIDISGFRQVNVHEKKNQDGTTFQKLKEEAFKLSGDEITELMRFSFKP  
CC  
EYTPITVTSADTSNNEILSKIQOQFETILTRENSTVNGTIEDPMGDKINLQNGQTLQPSDYTLQNDG  
cc  
SVMKDGATGGPNNDGILKGVKLEYIGNKLYVRGLNLEGQKVTLTYDVKLDDSFISNKFYDTNGRTTL  
cc  
NPKSEDPNTLRDFPIPKIRDVREYPTITIKNEKKGHEIFIKVDKDNKLLKLGATFELQEFNEDYKLYL  
CC  
PIKNNNSKVVTGENGKISYKDLKDGKYLIEAVSPEDYQKITNKPILTFEVKGSIKNIIAVNKQISEYH  
ecc  
EEGDKHLITNTHIPPKGIIPMTGGKILSFLIGGAMMSIAGGIYTWKRYKKSSDMSIKKD  
HhCC

Sequence length: 901

PHD :

Alpha helix	(Hh) :	148 is	16.43%	3 <sub>10</sub> helix	(Gg) :	0 is	0.00%
Pi helix	(Ii) :	0 is	0.00%	Beta bridge	(Bb) :	0 is	0.00%
Extended strand	(Ee) :	243 is	26.97%	Beta turn	(Tt) :	0 is	0.00%
Bend region	(Ss) :	0 is	0.00%	Random coil	(Cc) :	510 is	56.60%
Ambiguous states (?) :		0 is	0.00%	Other states		0 is	0.00%





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Figure 13

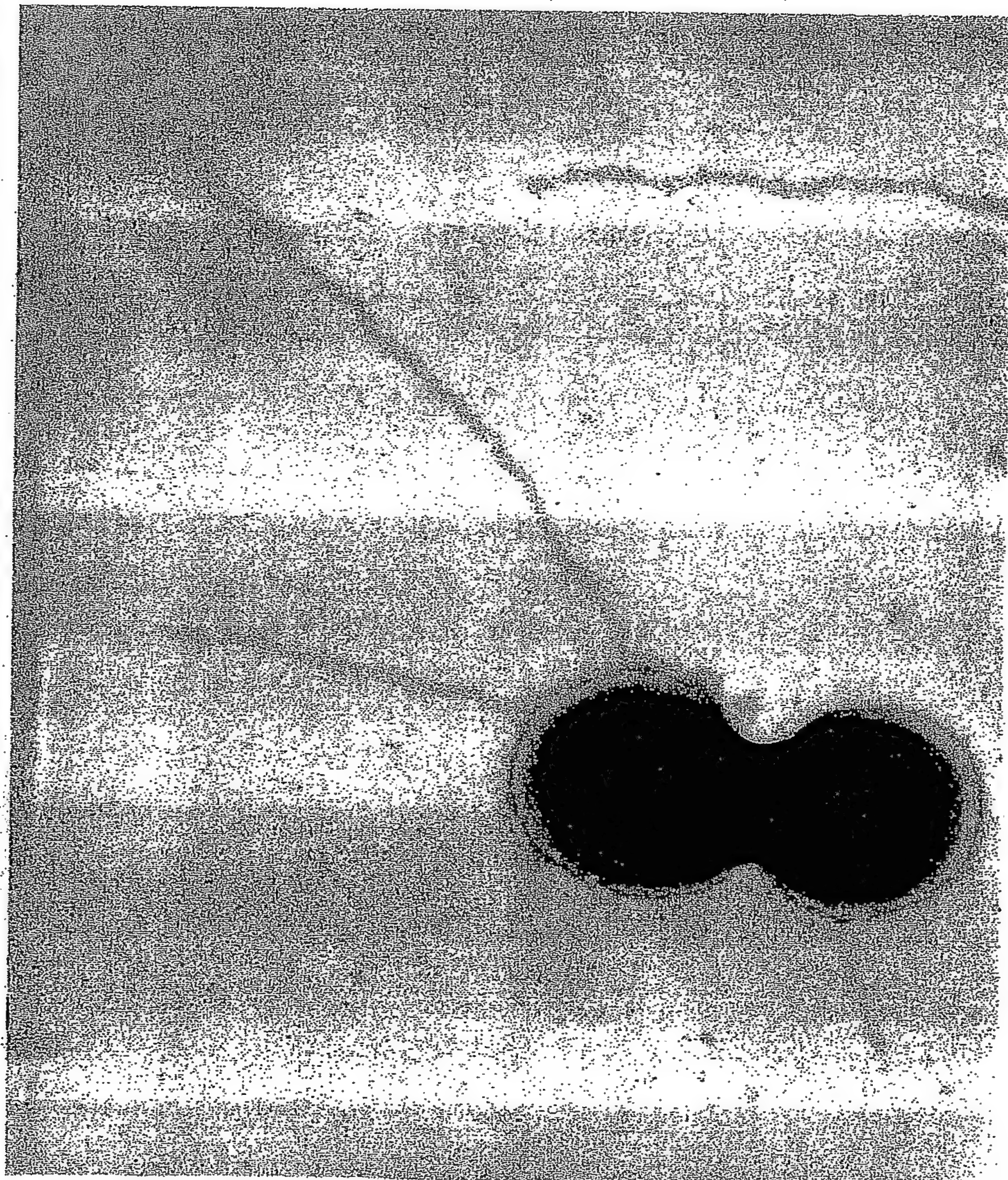
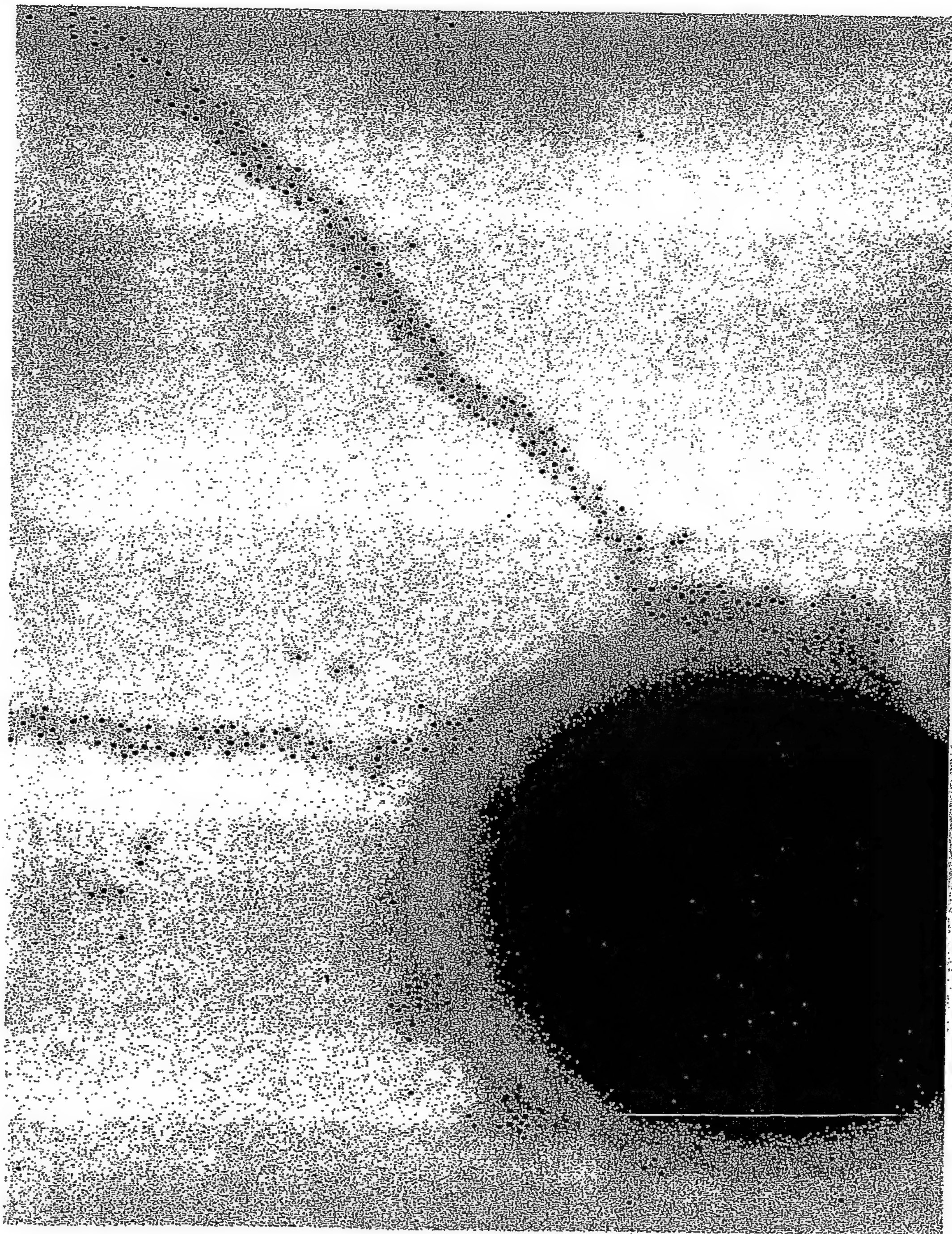




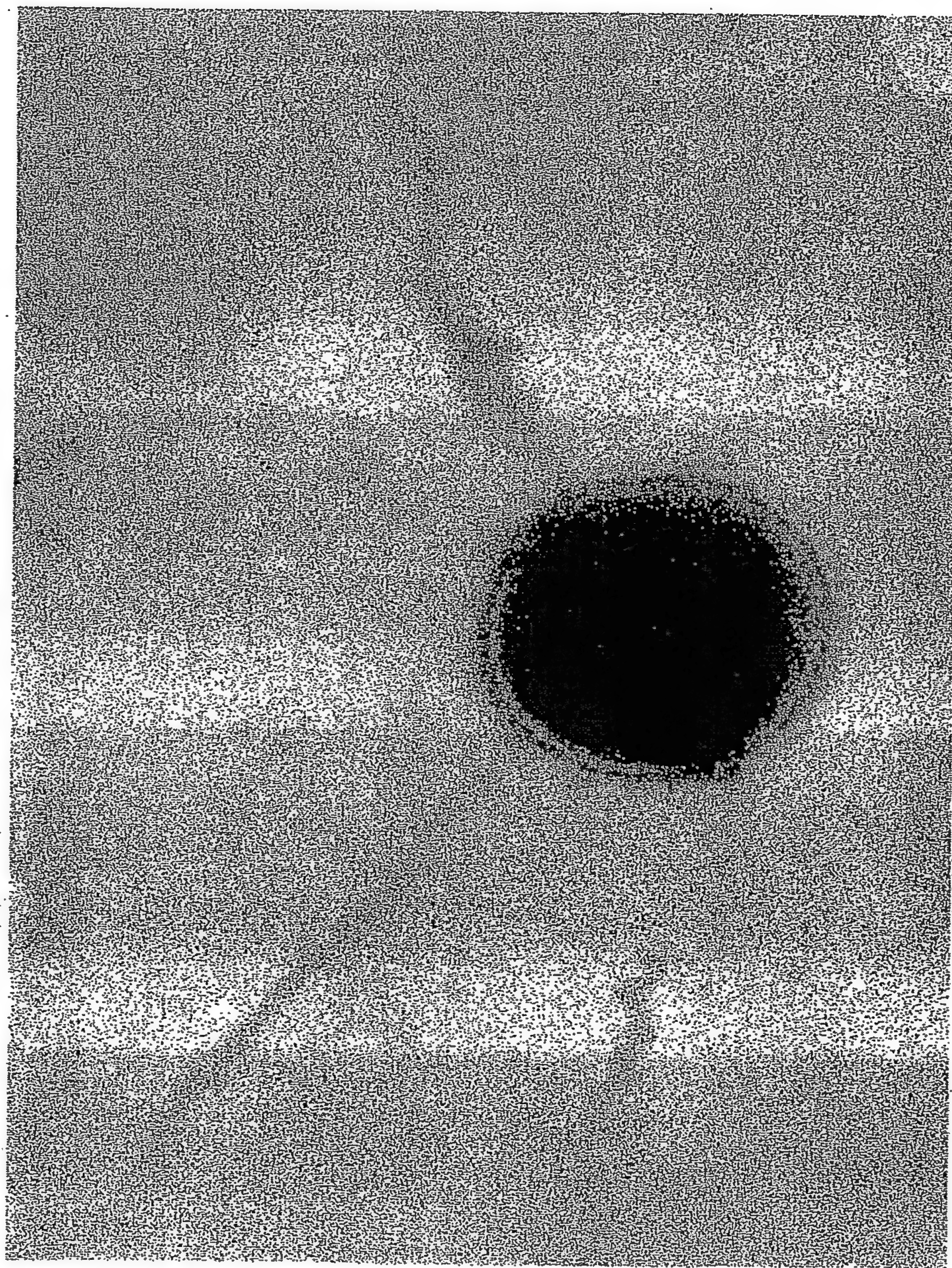
Figure 14





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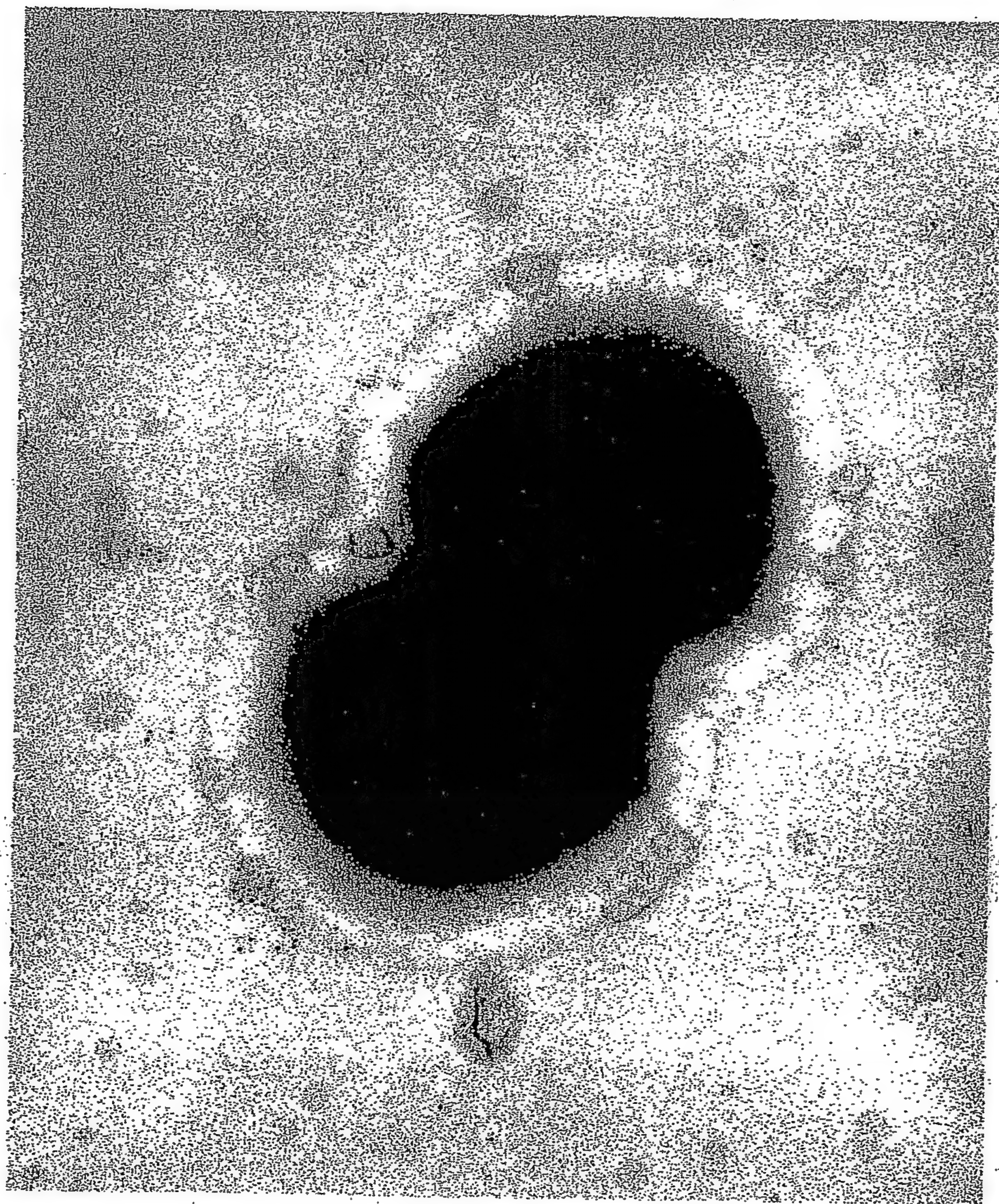
Figure 15





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Figure 16

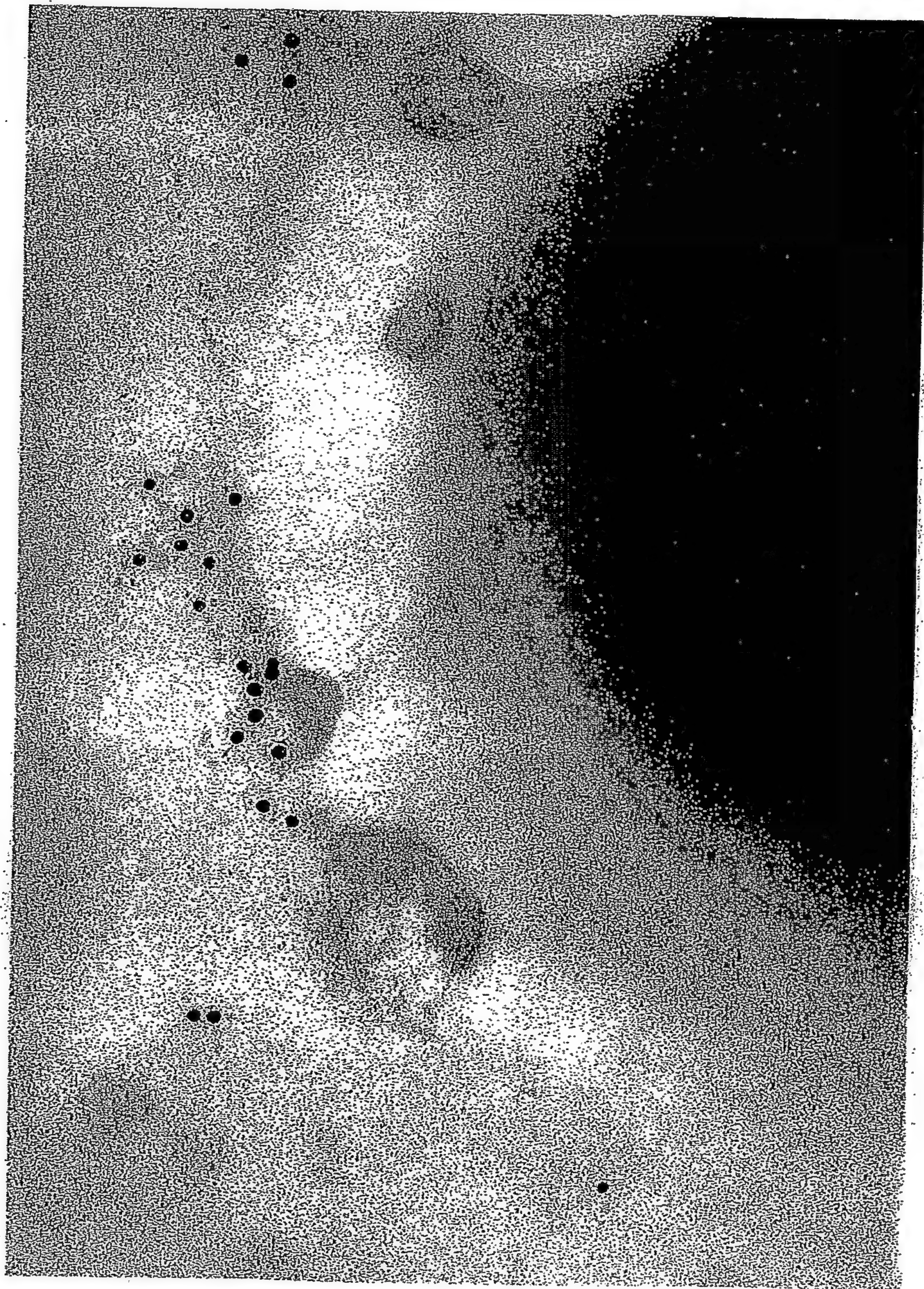




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Figure 17





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		T A A C A C G G A T T G G A A A T A A C A T C A A A T T G C G C G T G C T A T C A A C C A A	Majority
		10 20 30 40 50	
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1		T A A C A C G G A T T G G A A A T A A C A T C A A A T T G C G C G T G C T A T C A A C C A A	18rs21_all.seq
1		T A A C A C G G A T T G G A A A T A A C A T C A A A T T G C G C G T G C T A T C A A C C A A	cohl_all.seq
1		T A A C A C G G A T T G G A A A T A A C A T C A A A T T G C G C G T G C T A T C A A C C A A	cjb111_all.seq
1		T A A C A C G G A T T G G A A A T A A C A T C A A A T T G C G C G T G C T A T C A A C C A A	nen316_all.seq
1		T A A C A C G G A T T G G A A A T A A C A T C A A A T T G C G C G T G C T A T C A A C C A A	a909_all.seq
		A G C C A T C T A A T T T T C T C A G G C T C A G T A T C A C T T T C A T C G A G A A G C A C G T T A	Majority
		60 70 80 90 100	
51		A G C C A T C T A A T T T T C T C A G G C T C A G T A T C A C T T T C A T C G A G A A G C A C G T T A	2603_all.seq
1		A G C C A T C T A A T T T T C T C A G G C T C A G T A T C A C T T T C A T C G A G A A G C A C G T T A	18rs21_all.seq
51		A G C C A T C T A A T T T T C T C A G G C T C A G T A T C A C T T T C A T C G A G A A G C A C G T T A	cohl_all.seq
51		A G C C A T C T A A T T T T C T C A G G C T C A G T A T C A C T T T C A T C G A G A A G C A C G T T A	cjb111_all.seq
51		A G C C A T C T A A T T T T C T C A G G C T C A G T A T C A C T T T C A T C G A G A A G C A C G T T A	nen316_all.seq
51		A G C C A T C T A A T T T T C T C A G G C T C A G T A T C A C T T T C A T C G A G A A G C A C G T T A	a909_all.seq
		A G C C C A A C A G C A G G G G G G G T T G C T C T G A T T C G G T T A A G T A A T A A G C A A A A	Majority
		110 120 130 140 150	
101		A G C C C A A C A G C A G G G G G G G T T G C T C T G A T T C G G T T A A G T A A T A A G C A A A A	2603_all.seq
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101		A G C C C A A C A G C A G G G G G G G T T G C T C T G A T T C G G T T A A G T A A T A A G C A A A A	cjb111_all.seq
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101		A G C C C A A C A G C A G G G G G G G T T G C T C T G A T T C G G T T A A G T A A T A A G C A A A A	a909_all.seq
		T C C T C A C C A A T T T T T C C C A G T A A T A A G A G G C G T T G T T G A G G T A T A G G G A T T	Majority
		160 170 180 190 200	
150		T C C T C A C C A A T T T T T C C C A G T A A T A A G A G G C G T T G T T G A G G T A T A G G G A T T	2603_all.seq
1		T C C T C A C C A A T T T T T C C C A G T A A T A A G A G G C G T T G T T G A G G T A T A G G G A T T	18rs21_all.seq
151		T C C T C A C C A A T T T T T C C C A G T A A T A A G A G G C G T T G T T G A G G T A T A G G G A T T	cohl_all.seq
150		T C C T C A C C A A T T T T T C C C A G T A A T A A G A G G C G T T G T T G A G G T A T A G G G A T T	cjb111_all.seq
51		T C C T C A C C A A T T T T T C C C A G T A A T A A G A G G C G T T G T T G A G G T A T A G G G A T T	nen316_all.seq
51		T C C T C A C C A A T T T T T C C C A G T A A T A A G A G G C G T T G T T G A G G T A T A G G G A T T	a909_all.seq
		A C C T G T G C C A T A G T C G A T G A T G G T C A C A A A A T G A C C A T T C C C C A T A A A A G	Majority
		210 220 230 240 250	
200		A C C T G T G C C A T A G T C G A T G A T G G T C A C A A A A T G A C C A T T C C C C A T A A A A G	2603_all.seq
1		A C C T G T G C C A T A G T C G A T G A T G G T C A C A A A A T G A C C A T T C C C C A T A A A A G	18rs21_all.seq
201		A C C T G T G C C A T A G T C G A T G A T G G T C A C A A A A T G A C C A T T C C C C A T A A A A G	cohl_all.seq
200		A C C T G T G C C A T A G T C G A T G A T G G T C A C A A A A T G A C C A T T C C C C A T A A A A G	cjb111_all.seq
201		A C C T G T G C C A T A G T C G A T G A T G G T C A C A A A A T G A C C A T T C C C C A T A A A A G	nen316_all.seq
201		A C C T G T G C C A T A G T C G A T G A T G G T C A C A A A A T G A C C A T T C C C C A T A A A A G	a909_all.seq
		G G C C A A C C A A A A A C T T C A C C T G T G G C C G T T T T T T G A T G T C A A C A C C A G T A T	Majority
		260 270 280 290 300	
50		G G C C A A C C A A A A A C T T C A C C T G T G G C C G T T T T T T G A T G T C A A C A C C A G T A T	2603_all.seq
9		G G C C A A C C A A A A A C T T C A C C T G T G G C C G T T T T T T G A T G T C A A C A C C A G T A T	18rs21_all.seq
51		G G C C A A C C A A A A A C T T C A C C T G T G G C C G T T T T T T G A T G T C A A C A C C A G T A T	cohl_all.seq
50		G G C C A A C C A A A A A C T T C A C C T G T G G C C G T T T T T T G A T G T C A A C A C C A G T A T	cjb111_all.seq
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51		G G C C A A C C A A A A A C T T C A C C T G T G G C C G T T T T T T G A T G T C A A C A C C A G T A T	a909_all.seq
		T T T G G A T G T A C C C C T T A A C A T G C C C T T T G G T A T C T G C A A C A G A G A T A A T A	Majority
		310 320 330 340 350	
200		T T T G G A T G T A C C C C T T A A C A T G C C C T T T G G T A T C T G C A A C A G A G A T A A T A	2603_all.seq
9		T T T G G A T G T A C C C C T T A A C A T G C C C T T T G G T A T C T G C A A C A G A G A T A A T A	18rs21_all.seq
201		T T T G G A T G T A C C C C T T A A C A T G C C C T T T G G T A T C T G C A A C A G A G A T A A T A	cohl_all.seq
200		T T T G G A T G T A C C C C T T A A C A T G C C C T T T G G T A T C T G C A A C A G A G A T A A T A	cjb111_all.seq
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201		T T T G G A T G T A C C C C T T A A C A T G C C C T T T G G T A T C T G C A A C A G A G A T A A T A	a909_all.seq

Figure 18



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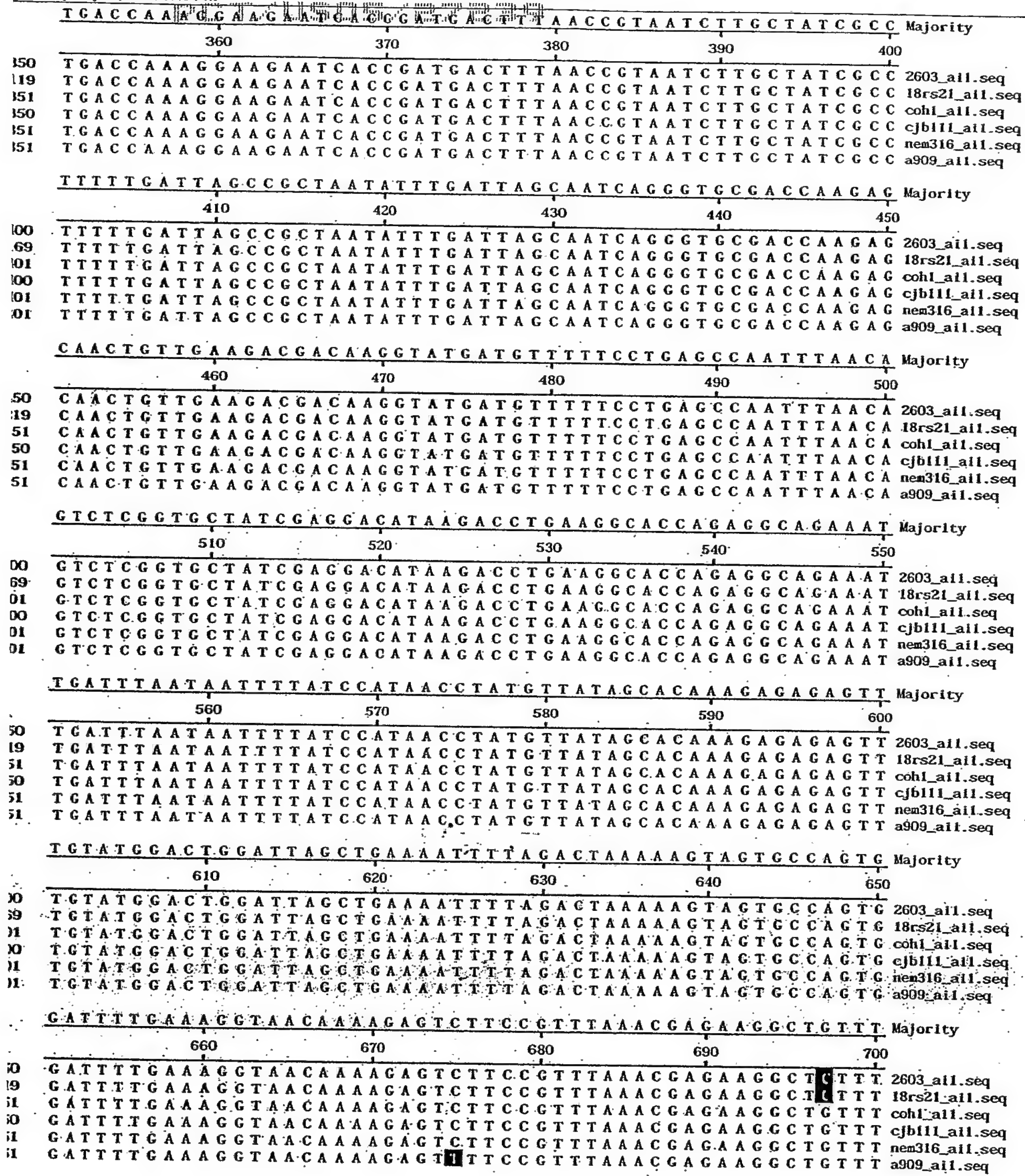


FIGURE 18 A

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710		720		730		740		750	
00	T	T	A	G	T	C	G	T	T
69	T	T	A	G	T	C	G	T	T
01	T	T	A	G	T	C	G	T	T
00	T	T	A	G	T	C	G	T	T
01	T	T	A	G	T	C	G	T	T
01	T	T	A	G	T	C	G	T	T
C A T G G A T T G C T A T G T T A C T G G C A T G A G G T C T C A C G A T A T T T A G T A A G A T A Majority									
760		770		780		790		800	
50	C	A	T	G	G	A	T	T	G
19	C	A	T	G	G	A	T	T	G
51	C	A	T	G	G	A	T	T	G
50	C	A	T	G	G	A	T	T	G
51	C	A	T	G	G	A	T	T	G
51	C	A	T	G	G	A	T	T	G
T T C G T T T G A A G A T A T T C C C A C G T A T T T T T T A A A G G T T T T A A G A A A A T G T G Majority									
810		820		830		840		850	
00	T	T	C	G	T	T	T	G	A
69	T	T	C	G	T	T	T	G	A
01	T	T	C	G	T	T	T	G	A
00	T	T	C	G	T	T	T	G	A
01	T	T	C	G	T	T	T	G	A
01	T	T	C	G	T	T	T	G	A
T G G T G T C G T A A A A A T G T A A T A A T T T C G C T A C T T C C C C A A T C G G T A C C C C T Majority									
860		870		880		890		900	
50	T	G	G	T	G	T	C	G	T
19	T	G	G	T	G	T	C	G	T
51	T	G	G	T	G	T	C	G	T
50	T	G	G	T	G	T	C	G	T
51	T	G	G	T	G	T	C	G	T
51	T	G	G	T	G	T	C	G	T
C T T T G C A G T A G T A A T T G T C C C T C C T T A A T T T T T G C C T T T A G A A T A T A A C T Majority									
910		920		930		940		950	
00	C	T	T	T	G	C	A	G	T
69	C	T	T	T	G	C	A	G	T
01	C	T	T	T	G	C	A	G	T
00	C	T	T	T	G	C	A	G	T
01	C	T	T	T	G	C	A	G	T
01	C	T	T	T	G	C	A	G	T
T T G C A A G G A A A T G T C A G A G T A T T T T T T A A A A A C T G A G C G T A A G T G G G A A T Majority									
960		970		980		990		1000	
00	T	T	G	C	A	A	G	G	A
9	T	T	G	C	A	A	G	G	A
01	T	T	G	C	A	A	G	G	A
00	T	T	G	C	A	A	G	G	A
01	T	T	G	C	A	A	G	G	A
01	T	T	G	C	A	A	G	G	A
C T G A G A T A T A T A G G T A G T T G G C A A T A T C A G A T A C T T T G A G T T T G G A G T A G Majority									
1010		1020		1030		1040		1050	
00	C	T	G	A	G	A	T	A	T
9	C	T	G	A	G	A	T	A	T
01	C	T	G	A	G	A	T	A	T
00	C	T	G	A	G	A	T	A	T
01	C	T	G	A	G	A	T	A	T
01	C	T	G	A	G	A	T	A	T

FIGURE 18 B



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WO 2006/078318

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Alignment Report of Al-1 alignment, using J. Hein method with Weighted residue weight table.  
Thursday, July 29, 2004 5:46 PM

Page 4

A G G T G G T T G T C C A C A T A A T G G A G A A T A C T A T T G T A C A T T T G C T G C T T G T C Majority																																																	
1060										1070										1080										1090										1100									
1050	A G G T G G T T G T C C A C A T A A T G G A G A A T A C T A T T G T A C A T T T G C T G C T T G T C																													2603_all.seq																			
819	A G G T G G T T G T C C A C A T A A T G G A G A A T A C T A T T G T A C A T T T G C T G C T T G T C																													18rs21_all.seq																			
1051	A G G T G G T T G T C C A C A T A A T G G A G A A T A C T A T T G T A C A T T T G C T G C T T G T C																													cohl_all.seq																			
1050	A G G T G G T T G T C C A C A T A A T G G A G A A T A C T A T T G T A C A T T T G C T G C T T G T C																													cjb111_all.seq																			
1051	A G G T G G T T G T C C A C A T A A T G G A G A A T A C T A T T G T A C A T T T G C T G C T T G T C																													nem316_all.seq																			
1051	A G G T G G T T G T C C A C A T A A T G G A G A A T A C T A T T G T A C A T T T G C T G C T T G T C																													a909_all.seq																			
A G A G A T G C T C T T A T T G G T T A A G G A T T C T G A A A A A T C A A T A A G A G C T G C A C Majority																																																	
1110										1120										1130										1140										1150									
1100	A G A G A T G C T C T T A T T G G T T A A G G A T T C T G A A A A A T C A A T A A G A G C T G C A C																													2603_all.seq																			
869	A G A G A T G C T C T T A T T G G T T A A G G A T T C T G A A A A A T C A A T A A G A G C T G C A C																													18rs21_all.seq																			
1101	A G A G A T G C T C T T A T T G G T T A A G G A T T C T G A A A A A T C A A T A A G A G C T G C A C																													cohl_all.seq																			
1100	A G A G A T G C T C T T A T T G G T T A A G G A T T C T G A A A A A T C A A T A A G A G C T G C A C																													cjb111_all.seq																			
1101	A G A G A T G C T C T T A T T G G T T A A G G A T T C T G A A A A A T C A A T A A G A G C T G C A C																													nem316_all.seq																			
1101	A G A G A T G C T C T T A T T G G T T A A G G A T T C T G A A A A A T C A A T A A G A G C T G C A C																													a909_all.seq																			
A G C G A A T T C T T G A A A C A T C A A T A A G A T C A G G A G C C T C T T C G T T T A A A G C C Majority																																																	
1160										1170										1180										1190										1200									
1150	A G C G A A T T C T T G A A A C A T C A A T A A G A T C A G G A G C C T C T T C G T T T A A A G C C																													2603_all.seq																			
919	A G C G A A T T C T T G A A A C A T C A A T A A G A T C A G G A G C C T C T T C G T T T A A A G C C																													18rs21_all.seq																			
1151	A G C G A A T T C T T G A A A C A T C A A T A A G A T C A G G A G C C T C T T C G T T T A A A G C C																													cohl_all.seq																			
1150	A G C G A A T T C T T G A A A C A T C A A T A A G A T C A G G A G C C T C T T C G T T T A A A G C C																													cjb111_all.seq																			
1151	A G C G A A T T C T T G A A A C A T C A A T A A G A T C A G G A G C C T A T T C G T T T A A A G C C																													nem316_all.seq																			
1151	A G C G A A T T C T T G A A A C A T C A A T A A G A T C A G G A G C C T C T T C G T T T A A A G C C																													a909_all.seq																			
A T A T A G T G C T T T A C C A G C G C A T A A C T T T T A G C C A C A T C A G T A T T T T C C T C Majority																																																	
1210										1220										1230										1240										1250									
1200	A T A T A G T G C T T T A C C A G C G C A T A A C T T T T A G C C A C A T C A G T A T T T T C C T C																													2603_all.seq																			
969	A T A T A G T G C T T T A C C A G C G C A T A A C T T T T A G C C A C A T C A G T A T T T T C C T C																													18rs21_all.seq																			
1201	A T A T A G T G C T T T A C C A G C G C A T A A C T T T T A G C C A C A T C A G T A T T T T C C T C																													cohl_all.seq																			
1200	A T A T A G T G C T T T A C C A G C G C A T A A C T T T T A G C C A C A T C A G T A T T T T C C T C																													cjb111_all.seq																			
1201	A T A T A G T G C T T T A C C A G C G C A T A A C T T T T A G C C A C A T C A G T A T T T T C C T C																													nem316_all.seq																			
1201	A T A T A G T G C T T T A C C A G C G C A T A A C T T T T A G C C A C A T C A G T A T T T T C C T C																													a909_all.seq																			
G A A A C T T A A T T C T A G T A A T T T T G T T A A G T A A A C A A C A G T T A A G T T C T T T T Majority																																																	
1260										1270										1280										1290										1300									
250	G A A A C T T A A T T C T A G T A A T T T T G T T A A G T A A A C A A C A G T T A A G T T C T T T T																													2603_all.seq																			
019	G A A A C T T A A T T C T A G T A A T T T T G T T A A G T A A A C A A C A G T T A A G T T C T T T T																													18rs21_all.seq																			
251	G A A A C T T A A T T C T A G T A A T T T T G T T A A G T A A A C A A C A G T T A A G T T C T T T T																													cohl_all.seq																			
250	G A A A C T T A A T T C T A G T A A T T T T G T T A A G T A A A C A A C A G T T A A G T T C T T T T																													cjb111_all.seq																			
251	G A A A C T T A A T T C T A G T A A T T T T G T T A A G T A A A C A A C A G T T A A G T T C T T T T																													nem316_all.seq																			
251	G A A A C T T A A T T C T A G T A A T T T T G T T A A G T A A A C A A C A G T T A A G T T C T T T T																													a909_all.seq																			
C A G C T C T T A G G G C A G G G A T T G A A G A T G A G G T A A C A C T G G A T G A T G G G A G G Majority																																																	
1310										1320										1330										1340										1350									
300	C A G C T C T T A G G G C A G G G A T T G A A G A T G A G G T A A C A C T G G A T G A T G G G A G G																													2603_all.seq																			
069	C A G C T C T T A G G G C A G G G A T T G A A G A T G A G G T A A C A C T G G A T G A T G G G A G G																													18rs21_all.seq																			
301	C A G C T C T T A G G G C A G G G A T T G A A G A T G A G G T A A C A C T G G A T G A T G G G A G G																													cohl_all.seq																			
300	C A G C T C T T A G G G C A G G G A T T G A A G A T G A G G T A A C A C T G G A T G A T G G G A G G																													cjb111_all.seq																			
301	C A G C T C T T A G G G C A G G G A T T G A A G A T G A G G T A A C A C T G G A T G A T G G G A G G																													nem316_all.seq																			
301	C A G C T C T T A G G G C A G G G A T T G A A G A T G A G G T A A C A C T G G A T G A T G G G A G G																													a909_all.seq																			
C G A T T A A T T T C T T G C T T T A A C A G T T C A G T G T T A C C C A G C T T A A C G A G A T C Majority																																																	
1360										1370										1380										1390										1400									
350	C G A T T A A T T T C T T G C T T T A A C A G T T C A G T G T T A C C C A G C T T A A C G A G A T C																													2603_all.seq																			
119	C G A T T A A T T T C T T G C T T T A A C A G T T C A G T G T T A C C C A G C T T A A C G A G A T C																													18rs21_all.seq																			
351	C G A T T A A T T T C T T G C T T T A A C A G T T C A G T G T T A C C C A G C T T A A C G A G A T C																													cohl_all.seq																			
350	C G A T T A A T T T C T T G C T T T A A C A G T T C A G T G T T A C C C A G C T T A A C G A G A T C																													cjb111_all.seq																			
351	C G A T T A A T T T C T T G C T T T A A C A G T T C A G T G T T A C C C A G C T T A A C G A G A T C																													nem316_all.seq																			
351	C G A T T A A T T T C T T G C T T T A A C A G T T C A G T G T T A C C C A G C T T A A C G A G A T C																													a909_all.seq																			

	Majority									
	AATAATGTCGATTGAGATGCTTTTAAACAGTGGGTAACTGAAAAGAGTTT									
	1410	1420	1430	1440	1450					
400	AATAATGTCGATTGAGATGCTTTTAAACAGTGGGTAACTGAAAAGAGTTT									
169	AATAATGTCGATTGAGATGCTTTTAAACAGTGGGTAACTGAAAAGAGTTT									
401	AATAATGTCGATTGAGATGCTTTTAAACAGTGGGTAACTGAAAAGAGTTT									
400	AATAATGTCGATTGAGATGCTTTTAAACAGTGGGTAACTGAAAAGAGTTT									
401	AATAATGTCGATTGAGATGCTTTTAAACAGTGGGTAACTGAAAAGAGTTT									
401	AATAATGTCGATTGAGATGCTTTTAAACAGTGGGTAACTGAAAAGAGTTT									
	TCTTAGTATGTTTTAGGTGAAGACAATATCAGGATCCGCAACAATCTGT									
	1460	1470	1480	1490	1500					
450	TCTTAGTATGTTTTAGGTGAAGACAATATCAGGATCCGCAACAATCTGT									
219	TCTTAGTATGTTTTAGGTGAAGACAATATCAGGATCCGCAACAATCTGT									
451	TCTTAGTATGTTTTAGGTGAAGACAATATCAGGATCCGCAACAATCTGT									
450	TCTTAGTATGTTTTAGGTGAAGACAATATCAGGATCCGCAACAATCTGT									
451	TCTTAGTATGTTTTAGGTGAAGACAATATCAGGATCCGCAACAATCTGT									
451	TCTTAGTATGTTTTAGGTGAAGACAATATCAGGATCCGCAACAATCTGT									
	TCTGACTCTTCTAATAAATGATTGATGGCTTGTGCGCAACTAGCCTCAAA									
	1510	1520	1530	1540	1550					
500	TCTGACTCTTCTAATAAATGATTGATGGCTTGTGCGCAACTAGCCTCAAA									
269	TCTGACTCTTCTAATAAATGATTGATGGCTTGTGCGCAACTAGCCTCAAA									
501	TCTGACTCTTCTAATAAATGATTGATGGCTTGTGCGCAACTAGCCTCAAA									
500	TCTGACTCTTCTAATAAATGATTGATGGCTTGTGCGCAACTAGCCTCAAA									
501	TCTGACTCTTCTAATAAATGATTGATGGCTTGTGCGCAACTAGCCTCAAA									
501	TCTGACTCTTCTAATAAATGATTGATGGCTTGTGCGCAACTAGCCTCAAA									
	CTGTGTTTGGAAAAAGGCATCGATAGACACAAGAAGACTACGTATACTGG									
	1560	1570	1580	1590	1600					
550	CTGTGTTTGGAAAAAGGCATCGATAGACACAAGAAGACTACGTATACTGG									
319	CTGTGTTTGGAAAAAGGCATCGATAGACACAAGAAGACTACGTATACTGG									
551	CTGTGTTTGGAAAAAGGCATCGATAGACACAAGAAGACTACGTATACTGG									
550	CTGTGTTTGGAAAAAGGCATCGATAGACACAAGAAGACTACGTATACTGG									
551	CTGTGTTTGGAAAAAGGCATCGATAGACACAAGAAGACTACGTATACTGG									
551	CTGTGTTTGGAAAAAGGCATCGATAGACACAAGAAGACTACGTATACTGG									
	TAGTAGGAAAACAAGGGACAAGCTTTATATAGGATAAGATTTCTTTTTTA									
	1610	1620	1630	1640	1650					
600	TAGTAGGAAAACAAGGGACAAGCTTTATATAGGATAAGATTTCTTTTTTA									
169	TAGTAGGAAAACAAGGGACAAGCTTTATATAGGATAAGATTTCTTTTTTA									
601	TAGTAGGAAAACAAGGGACAAGCTTTATATAGGATAAGATTTCTTTTTTA									
600	TAGTAGGAAAACAAGGGACAAGCTTTATATAGGATAAGATTTCTTTTTTA									
601	TAGTAGGAAAACAAGGGACAAGCTTTATATAGGATAAGATTTCTTTTTTA									
601	TAGTAGGAAAACAAGGGACAAGCTTTATATAGGATAAGATTTCTTTTTTA									
	CTACGATGAGAAAATTGTTCTAGAAAGCGACTGGATAACTGTTCTTGCCCT									
	1660	1670	1680	1690	1700					
650	CTACGATGAGAAAATTGTTCTAGAAAGCGACTGGATAACTGTTCTTGCCCT									
119	CTACGATGAGAAAATTGTTCTAGAAAGCGACTGGATAACTGTTCTTGCCCT									
651	CTACGATGAGAAAATTGTTCTAGAAAGCGACTGGATAACTGTTCTTGCCCT									
650	CTACGATGAGAAAATTGTTCTAGAAAGCGACTGGATAACTGTTCTTGCCCT									
651	CTACGATGAGAAAATTGTTCTAGAAAGCGACTGGATAACTGTTCTTGCCCT									
651	CTACGATGAGAAAATTGTTCTAGAAAGCGACTGGATAACTGTTCTTGCCCT									
	ATTGATATCAGGGCTATAGGGATAAAATGGTCCAATAGCAATAAGATATT									
	1710	1720	1730	1740	1750					
700	ATTGATATCAGGGCTATAGGGATAAAATGGTCCAATAGCAATAAGATATT									
69	ATTGATATCAGGGCTATAGGGATAAAATGGTCCAATAGCAATAAGATATT									
701	ATTGATATCAGGGCTATAGGGATAAAATGGTCCAATAGCAATAAGATATT									
700	ATTGATATCAGGGCTATAGGGATAAAATGGTCCAATAGCAATAAGATATT									
701	ATTGATATCAGGGCTATAGGGATAAAATGGTCCAATAGCAATAAGATATT									
701	ATTGATATCAGGGCTATAGGGATAAAATGGTCCAATAGCAATAAGATATT									

**FIGURE 18 D**



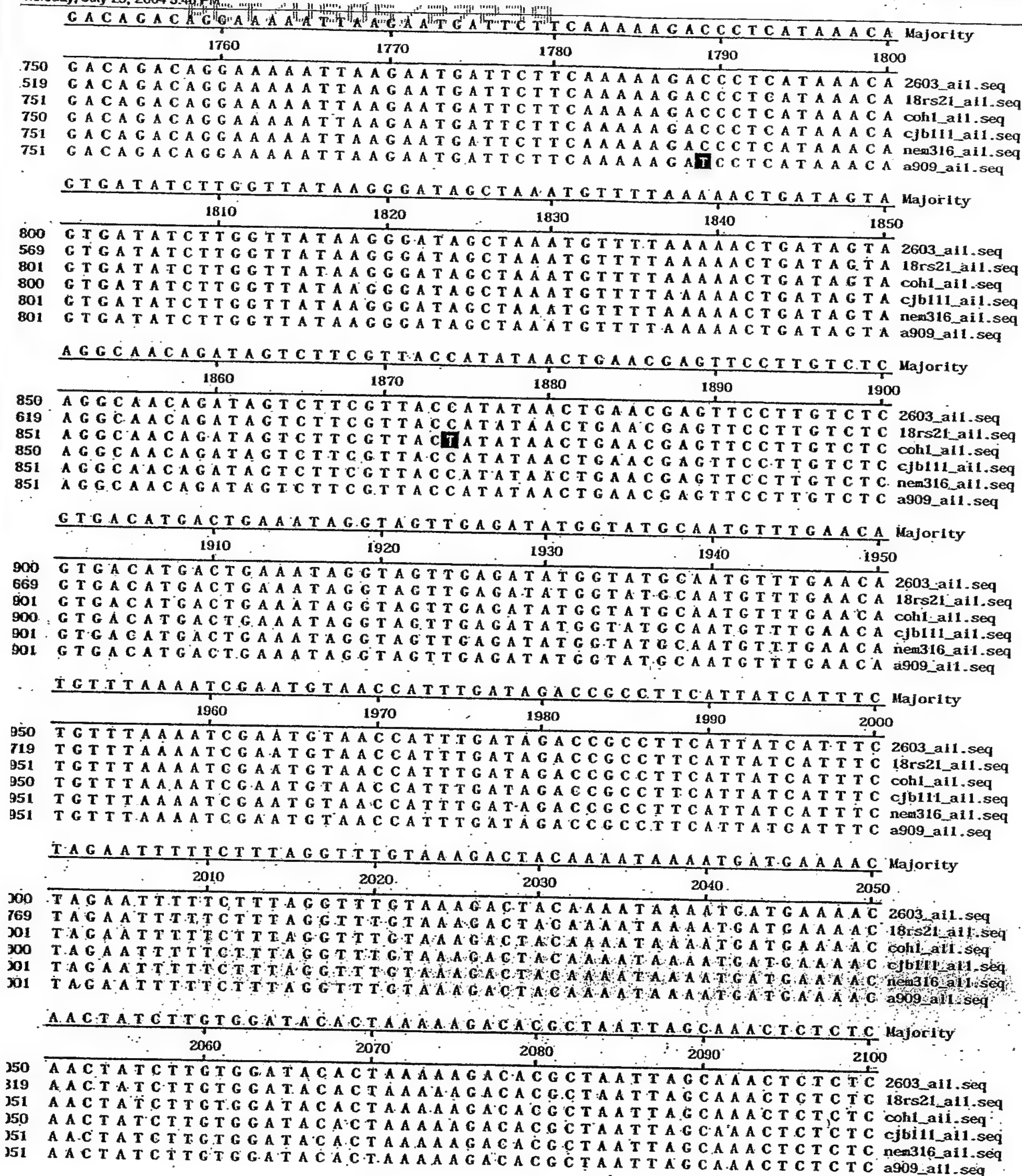


FIGURE 18 E

TTCATCACTCTCTCACCATTATTATACTACTATTTATATGACAAATAAAGG Majority									
2110		2120		2130		2140		2150	
2100	TTCATCACTCTCTCACCATTATTATACTACTATTTATATGACAAATAAAGG								2603_all.seq
1869	TTCATCACTCTCTCACCATTATTATACTACTATTTATATGACAAATAAAGG								18rs21_all.seq
2101	TTCATCACTCTCTCACCATTATTATACTACTATTTATATGACAAATAAAGG								coh1_all.seq
2100	TTCATCACTCTCTCACCATTATTATACTACTATTTATATGACAAATAAAGG								cjb111_all.seq
2101	TTCATCACTCTCTCACCATTATTATACTACTATTTATATGACAAATAAAGG								nem316_all.seq
2101	TTCATCACTCTCTCACCATTATTATACTACTATTTATATGACAAATAAAGG								a909_all.seq
TGATTTTGTAAATAATAACTTTGAAAATCCACATATATTTTTTAATCTT Majority									
2160		2170		2180		2190		2200	
2150	TGATTTTGTAAATAATAACTTTGAAAATCCACATATATTTTTTTAATCTT								2603_all.seq
1919	TGATTTTGTAAATAATAACTTTGAAAATCCACATATATTTTTTTAATCTT								18rs21_all.seq
2151	TGATTTTGTAAATAATAACTTTGAAAATCCACATATATTTTTTTAATCTT								coh1_all.seq
2150	TGATTTTGTAAATAATAACTTTGAAAATCCACATATATTTTTTTAATCTT								cjb111_all.seq
2151	TGATTTTGTAAATAATAACTTTGAAAATCCACATATATTTTTTTAATCTT								nem316_all.seq
2151	TGATTTTGTAAATAATAACTTTGAAAATCCACATATATTTTTTTAATCTT								a909_all.seq
CCGTCTGAAAAA-TAAATAAAAAATAGTAAAAATAAACACGAATTTAAAAA Majority									
2210		2220		2230		2240		2250	
2199	CCGTCTGAAAAA-TAAATAAAAAATAGTAAAAATAAACACGAATTTAAAAA								2603_all.seq
1968	CCGTCTGAAAAA-TAAATAAAAAATAGTAAAAATAAACACGAATTTAAAAA								18rs21_all.seq
2200	CCGTCTGAAAAA-TAAATAAAAAATAGTAAAAATAAACACGAATTTAAAAA								coh1_all.seq
2200	CCGTCTGAAAAA-TAAATAAAAAATAGTAAAAATAAACACGAATTTAAAAA								cjb111_all.seq
2201	CCGTCTGAAAAA-TAAATAAAAAATAGTAAAAATAAACACGAATTTAAAAA								nem316_all.seq
2201	CCGTCTGAAAAA-TAAATAAAAAATAGTAAAAATAAACACGAATTTAAAAA								a909_all.seq
TAAGCAAATTTTTTTAAGAAAATCTGTGCTAAACTTTAATAGTTTTGTGCT Majority									
2260		2270		2280		2290		2300	
2248	TAAGCAAATTTTTTTAAGAAAATCTGTGCTAAACTTTAATAGTTTTGTGCT								2603_all.seq
2017	TAAGCAAATTTTTTTAAGAAAATCTGTGCTAAACTTTAATAGTTTTGTGCT								18rs21_all.seq
2249	TAAGCAAATTTTTTTAAGAAAATCTGTGCTAAACTTTAATAGTTTTGTGCT								coh1_all.seq
2249	TAAGCAAATTTTTTTAAGAAAATCTGTGCTAAACTTTAATAGTTTTGTGCT								cjb111_all.seq
2251	TAAGCAAATTTTTTTAAGAAAATCTGTGCTAAACTTTAATAGTTTTGTGCT								nem316_all.seq
2250	TAAGCAAATTTTTTTAAGAAAATCTGTGCTAAACTTTAATAGTTTTGTGCT								a909_all.seq
TAATAATAATCAGCACTTACAAAGAACAAAGGGAAAAGCGAGGAGAGAAAC Majority									
2310		2320		2330		2340		2350	
2298	TAATAATAATCAGCACTTACAAAGAACAAAGGGAAAAGCGAGGAGAGAAAC								2603_all.seq
2067	TAATAATAATCAGCACTTACAAAGAACAAAGGGAAAAGCGAGGAGAGAAAC								18rs21_all.seq
2299	TAATAATAATCAGCACTTACAAAGAACAAAGGGAAAAGCGAGGAGAGAAAC								coh1_all.seq
2299	TAATAATAATCAGCACTTACAAAGAACAAAGGGAAAAGCGAGGAGAGAAAC								cjb111_all.seq
301	TAATAATAATCAGCACTTACAAAGAACAAAGGGAAAAGCGAGGAGAGAAAC								nem316_all.seq
300	TAATAATAATCAGCACTTACAAAGAACAAAGGGAAAAGCGAGGAGAGAAAC								a909_all.seq
TTTTAATGAAATTATCGAAGAAAGTTATTGTTTTCGGCTGCTGTTTTTAACA Majority									
2360		2370		2380		2390		2400	
348	TTTTAATGAAATTATCGAAGAAAGTTATTGTTTTCGGCTGCTGTTTTTAACA								2603_all.seq
117	TTTTAATGAAATTATCGAAGAAAGTTATTGTTTTCGGCTGCTGTTTTTAACA								18rs21_all.seq
349	TTTTAATGAAATTATCGAAGAAAGTTATTGTTTTCGGCTGCTGTTTTTAACA								coh1_all.seq
349	TTTTAATGAAATTATCGAAGAAAGTTATTGTTTTCGGCTGCTGTTTTTAACA								cjb111_all.seq
351	TTTTAATGAAATTATCGAAGAAAGTTATTGTTTTCGGCTGCTGTTTTTAACA								nem316_all.seq
350	TTTTAATGAAATTATCGAAGAAAGTTATTGTTTTCGGCTGCTGTTTTTAACA								a909_all.seq
ATGGTGGCGGGGTCAACTGTTGAACCACTAGCTCAGTTTGGCGACTGGAAT Majority									
2410		2420		2430		2440		2450	
398	ATGGTGGCGGGGTCAACTGTTGAACCACTAGCTCAGTTTGGCGACTGGAAT								2603_all.seq
167	ATGGTGGCGGGGTCAACTGTTGAACCACTAGCTCAGTTTGGCGACTGGAAT								18rs21_all.seq
399	ATGGTGGCGGGGTCAACTGTTGAACCACTAGCTCAGTTTGGCGACTGGAAT								coh1_all.seq
399	ATGGTGGCGGGGTCAACTGTTGAACCACTAGCTCAGTTTGGCGACTGGAAT								cjb111_all.seq
401	ATGGTGGCGGGGTCAACTGTTGAACCACTAGCTCAGTTTGGCGACTGGAAT								nem316_all.seq
400	ATGGTGGCGGGGTCAACTGTTGAACCACTAGCTCAGTTTGGCGACTGGAAT								a909_all.seq

FIGURE 18 F



G A G T A T T G T A A G A G C T G C A G A A G T G T C A C A A G A A C G C C C A G C G A A A A C A A Majority									
2460		2470		2480		2490		2500	
2448	G A G T A T T G T A A G A G C T G C A G A A G T G T C A C A A G A A C G C C C A G C G A A A A C A A 2603_all.seq								
2217	G A G T A T T G T A A G A G C T G C A G A A G T G T C A C A A G A A C G C C C A G C G A A A A C A A 18rs21_all.seq								
2449	G A G T A T T G T A A G A G C T G C A G A A G T G T C A C A A G A A C G C C C A G C G A A A A C A A coh1_all.seq								
2449	G A G T A T T G T A A G A G C T G C A G A A G T G T C A C A A G A A C G C C C A G C G A A A A C A A cjb111_all.seq								
2451	G A G T A T T G T A A G A G C T G C A G A A G T G T C A C A A G A A C G C C C A G C G A A A A C A A nem316_all.seq								
2450	G A G T A T T G T A A G A G C T G C A G A A G T G T C A C A A G A A C G C C C A G C G A A A A C A A a909_all.seq								
C A G T A A A T A T C T A T A A A T T A C A A G C T G A T A G T T A T A A A T C G G A A A T T A C T Majority									
2510		2520		2530		2540		2550	
2498	C A G T A A A T A T C T A T A A A T T A C A A G C T G A T A G T T A T A A A T C G G A A A T T A C T 2603_all.seq								
2267	C A G T A A A T A T C T A T A A A T T A C A A G C T G A T A G T T A T A A A T C G G A A A T T A C T 18rs21_all.seq								
2499	C A G T A A A T A T C T A T A A A T T A C A A G C T G A T A G T T A T A A A T C G G A A A T T A C T coh1_all.seq								
2499	C A G T A A A T A T C T A T A A A T T A C A A G C T G A T A G T T A T A A A T C G G A A A T T A C T cjb111_all.seq								
2501	C A G T A A A T A T C T A T A A A T T A C A A G C T G A T A G T T A T A A A T C G G A A A T T A C T nem316_all.seq								
2500	C A G T A A A T A T C T A T A A A T T A C A A G C T G A T A G T T A T A A A T C G G A A A T T A C T a909_all.seq								
T C T A A T G G T G G T A T C G A G A A T A A A G A C G G C C A A G T A A T A T C T A A C T A T G C Majority									
2560		2570		2580		2590		2600	
2548	T C T A A T G G T G G T A T C G A G A A T A A A G A C G G C C A A G T A A T A T C T A A C T A T G C 2603_all.seq								
2317	T C T A A T G G T G G T A T C G A G A A T A A A G A C G G C C A A G T A A T A T C T A A C T A T G C 18rs21_all.seq								
2549	T C T A A T G G T G G T A T C G A G A A T A A A G A C G G C C A A G T A A T A T C T A A C T A T G C coh1_all.seq								
2549	T C T A A T G G T G G T A T C G A G A A T A A A G A C G G C C A A G T A A T A T C T A A C T A T G C cjb111_all.seq								
2551	T C T A A T G G T G G T A T C G A G A A T A A A G A C G G C C A A G T A A T A T C T A A C T A T G C nem316_all.seq								
2550	T C T A A T G G T G G T A T C G A G A A T A A A G A C G G C C A A G T A A T A T C T A A C T A T G C a909_all.seq								
T A A A C T T G G T G A C A A T G T A A A A G G T T T G C A A G G T G T A C A G T T T A A A C G T T Majority									
2610		2620		2630		2640		2650	
2598	T A A A C T T G G T G A C A A T G T A A A A G G T T T G C A A G G T G T A C A G T T T A A A C G T T 2603_all.seq								
2367	T A A A C T T G G T G A C A A T G T A A A A G G T T T G C A A G G T G T A C A G T T T A A A C G T T 18rs21_all.seq								
2599	T A A A C T T G G T G A C A A T G T A A A A G G T T T G C A A G G T G T A C A G T T T A A A C G T T coh1_all.seq								
2599	T A A A C T T G G T G A C A A T G T A A A A G G T T T G C A A G G T G T A C A G T T T A A A C G T T cjb111_all.seq								
2601	T A A A C T T G G T G A C A A T G T A A A A G G T T T G C A A G G T G T A C A G T T T A A A C G T T nem316_all.seq								
2600	T A A A C T T G G T G A C A A T G T A A A A G G T T T G C A A G G T G T A C A G T T T A A A C G T T a909_all.seq								
A T A A A G T C A A G A C G G A T A T T T C T G T T G A T G A A T T G A A A A A A T T G A C A A C A Majority									
2660		2670		2680		2690		2700	
2648	A T A A A G T C A A G A C G G A T A T T T C T G T T G A T G A A T T G A A A A A A T T G A C A A C A 2603_all.seq								
2417	A T A A A G T C A A G A C G G A T A T T T C T G T T G A T G A A T T G A A A A A A T T G A C A A C A 18rs21_all.seq								
2649	A T A A A G T C A A G A C G G A T A T T T C T G T T G A T G A A T T G A A A A A A T T G A C A A C A coh1_all.seq								
2649	A T A A A G T C A A G A C G G A T A T T T C T G T T G A T G A A T T G A A A A A A T T G A C A A C A cjb111_all.seq								
2651	A T A A A G T C A A G A C G G A T A T T T C T G T T G A T G A A T T G A A A A A A T T G A C A A C A nem316_all.seq								
2650	A T A A A G T C A A G A C G G A T A T T T C T G T T G A T G A A T T G A A A A A A T T G A C A A C A a909_all.seq								
G T T G A A G C A G C A G A T G C A A A A G T T G G A A C G A T T C T T G A A G A A G G T G T C A G Majority									
2710		2720		2730		2740		2750	
2698	G T T G A A G C A G C A G A T G C A A A A G T T G G A A C G A T T C T T G A A G A A G G T G T C A G 2603_all.seq								
2667	G T T G A A G C A G C A G A T G C A A A A G T T G G A A C G A T T C T T G A A G A A G G T G T C A G 18rs21_all.seq								
2699	G T T G A A G C A G C A G A T G C A A A A G T T G G A A C G A T T C T T G A A G A A G G T G T C A G coh1_all.seq								
2699	G T T G A A G C A G C A G A T G C A A A A G T T G G A A C G A T T C T T G A A G A A G G T G T C A G cjb111_all.seq								
2701	G T T G A A G C A G C A G A T G C A A A A G T T G G A A C G A T T C T T G A A G A A G G T G T C A G nem316_all.seq								
2700	G T T G A A G C A G C A G A T G C A A A A G T T G G A A C G A T T C T T G A A G A A G G T G T C A G a909_all.seq								
T C T A C C T C A A A A A A C T A A T G C T C A A G G T T T G G T C G T C G A T G C T C T G G A T T Majority									
2760		2770		2780		2790		2800	
2748	T C T A C C T C A A A A A A C T A A T G C T C A A G G T T T G G T C G T C G A T G C T C T G G A T T 2603_all.seq								
2717	T C T A C C T C A A A A A A C T A A T G C T C A A G G T T T G G T C G T C G A T G C T C T G G A T T 18rs21_all.seq								
2749	T C T A C C T C A A A A A A C T A A T G C T C A A G G T T T G G T C G T C G A T G C T C T G G A T T coh1_all.seq								
2749	T C T A C C T C A A A A A A C T A A T G C T C A A G G T T T G G T C G T C G A T G C T C T G G A T T cjb111_all.seq								
2751	T C T A C C T C A A A A A A C T A A T G C T C A A G G T T T G G T C G T C G A T G C T C T G G A T T nem316_all.seq								
2750	T C T A C C T C A A A A A A C T A A T G C T C A A G G T T T G G T C G T C G A T G C T C T G G A T T a909_all.seq								

FIGURE 18 G

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C A A A A A G C T A A T G T G A G A T A C T T G T A T G T A G A A G A T T T A A A G A A T T C A C C T										Majority
2810		2820		2830		2840		2850		
2798	C A A A A A G T A A T G T G A G A T A C T T G T A T G T A G A A G A T T T A A A G A A T T C A C C T									2603_all.seq
2567	C A A A A A G T A A T G T G A G A T A C T T G T A T G T A G A A G A T T T A A A G A A T T C A C C T									18rs21_all.seq
2799	C A A A A A G T A A T G T G A G A T A C T T G T A T G T A G A A G A T T T A A A G A A T T C A C C T									cohl_all.seq
2799	C A A A A A G T A A T G T G A G A T A C T T G T A T G T A G A A G A T T T A A A G A A T T C A C C T									cjb111_all.seq
2801	C A A A A A G T A A T G T G A G A T A C T T G T A T G T A G A A G A T T T A A A G A A T T C A C C T									nem316_all.seq
2800	C A A A A A G T A A T G T G A G A T A C T T G T A T G T A G A A G A T T T A A A G A A T T C A C C T									a909_all.seq
T C A A A C A T T A C C A A A G C T T A T G C T G T A C C G T T T G T G T T G G A A T T A C C A G T										Majority
2860		2870		2880		2890		2900		
2848	T C A A A C A T T A C C A A A G C T T A T G C T G T A C C G T T T G T G T T G G A A T T A C C A G T									2603_all.seq
2617	T C A A A C A T T A C C A A A G C T T A T G C T G T A C C G T T T G T G T T G G A A T T A C C A G T									18rs21_all.seq
2849	T C A A A C A T T A C C A A A G C T T A T G C T G T A C C G T T T G T G T T G G A A T T A C C A G T									cohl_all.seq
2849	T C A A A C A T T A C C A A A G C T T A T G C T G T A C C G T T T G T G T T G G A A T T A C C A G T									cjb111_all.seq
2851	T C A A A C A T T A C C A A A G C T T A T G C T G T A C C G T T T G T G T T G G A A T T A C C A G T									nem316_all.seq
2850	T C A A A C A T T A C C A A A G C T T A T G C T G T A C C G T T T G T G T T G G A A T T A C C A G T									a909_all.seq
T G C T A A C T C T A C A G G T A C A G G T T T C C T T T C T G A A A T T A A T A T T T A C C C T A										Majority
2910		2920		2930		2940		2950		
2898	T G C T A A C T C T A C A G G T A C A G G T T T C C T T T C T G A A A T T A A T A T T T A C C C T A									2603_all.seq
2667	T G C T A A C T C T A C A G G T A C A G G T T T C C T T T C T G A A A T T A A T A T T T A C C C T A									18rs21_all.seq
2899	T G C T A A C T C T A C A G G T A C A G G T T T C C T T T C T G A A A T T A A T A T T T A C C C T A									cohl_all.seq
2899	T G C T A A C T C T A C A G G T A C A G G T T T C C T T T C T G A A A T T A A T A T T T A C C C T A									cjb111_all.seq
2901	T G C T A A C T C T A C A G G T A C A G G T T T C C T T T C T G A A A T T A A T A T T T A C C C T A									nem316_all.seq
2900	T G C T A A C T C T A C A G G T A C A G G T T T C C T T T C T G A A A T T A A T A T T T A C C C T A									a909_all.seq
A A A A C G T T G T A A C T G A T G A A C C A A A A A C A G A T A A A G A T G T T A A A A A A T T A										Majority
2960		2970		2980		2990		3000		
2948	A A A A C G T T G T A A C T G A T G A A C C A A A A A C A G A T A A A G A T G T T A A A A A A T T A									2603_all.seq
2717	A A A A C G T T G T A A C T G A T G A A C C A A A A A C A G A T A A A G A T G T T A A A A A A T T A									18rs21_all.seq
2949	A A A A C G T T G T A A C T G A T G A A C C A A A A A C A G A T A A A G A T G T T A A A A A A T T A									cohl_all.seq
2949	A A A A C G T T G T A A C T G A T G A A C C A A A A A C A G A T A A A G A T G T T A A A A A A T T A									cjb111_all.seq
2951	A A A A C G T T G T A A C T G A T G A A C C A A A A A C A G A T A A A G A T G T T A A A A A A T T A									nem316_all.seq
2950	A A A A C G T T G T A A C T G A T G A A C C A A A A A C A G A T A A A G A T G T T A A A A A A T T A									a909_all.seq
G G T C A G G A C C A T G C A G G T T A T A C G A T T G G T G A A G A A T T C A A A T G G T T C T T										Majority
3010		3020		3030		3040		3050		
998	G G T C A G G A C C A T G C A G G T T A T A C G A T T G G T G A A G A A T T C A A A T G G T T C T T									2603_all.seq
767	G G T C A G G A C C A T G C A G G T T A T A C G A T T G G T G A A G A A T T C A A A T G G T T C T T									18rs21_all.seq
999	G G T C A G G A C C A T G C A G G T T A T A C G A T T G G T G A A G A A T T C A A A T G G T T C T T									cohl_all.seq
999	G G T C A G G A C C A T G C A G G T T A T A C G A T T G G T G A A G A A T T C A A A T G G T T C T T									cjb111_all.seq
001	G G T C A G G A C C A T G C A G G T T A T A C G A T T G G T G A A G A A T T C A A A T G G T T C T T									nem316_all.seq
000	G G T C A G G A C C A T G C A G G T T A T A C G A T T G G T G A A G A A T T C A A A T G G T T C T T									a909_all.seq
G A A A T C T A C A A T C C C T G C C A A T T T A G G T G A C T A T G A A A A A T T T T G A A A T T A										Majority
3060		3070		3080		3090		3100		
048	G A A A T C T A C A A T C C C T G C C A A T T T A G G T G A C T A T G A A A A A T T T T G A A A T T A									2603_all.seq
817	G A A A T C T A C A A T C C C T G C C A A T T T A G G T G A C T A T G A A A A A T T T T G A A A T T A									18rs21_all.seq
049	G A A A T C T A C A A T C C C T G C C A A T T T A G G T G A C T A T G A A A A A T T T T G A A A T T A									cohl_all.seq
249	G A A A T C T A C A A T C C C T G C C A A T T T A G G T G A C T A T G A A A A A T T T T G A A A T T A									cjb111_all.seq
251	G A A A T C T A C A A T C C C T G C C A A T T T A G G T G A C T A T G A A A A A T T T T G A A A T T A									nem316_all.seq
250	G A A A T C T A C A A T C C C T G C C A A T T T A G G T G A C T A T G A A A A A T T T T G A A A T T A									a909_all.seq
C T G A T A A A T T T T G C A G A T G G C T T G A C T T A T A A A T C T G T T G G A A A A A T C A A G										Majority
3110		3120		3130		3140		3150		
298	C T G A T A A A T T T T G C A G A T G G C T T G A C T T A T A A A T C T G T T G G A A A A A T C A A G									2603_all.seq
367	C T G A T A A A T T T T G C A G A T G G C T T G A C T T A T A A A T C T G T T G G A A A A A T C A A G									18rs21_all.seq
299	C T G A T A A A T T T T G C A G A T G G C T T G A C T T A T A A A T C T G T T G G A A A A A T C A A G									cohl_all.seq
299	C T G A T A A A T T T T G C A G A T G G C T T G A C T T A T A A A T C T G T T G G A A A A A T C A A G									cjb111_all.seq
101	C T G A T A A A T T T T G C A G A T G G C T T G A C T T A T A A A T C T G T T G G A A A A A T C A A G									nem316_all.seq
100	C T G A T A A A T T T T G C A G A T G G C T T G A C T T A T A A A T C T G T T G G A A A A A T C A A G									a909_all.seq

FIGURE 18 H



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A T T G G T T C G A A A A C A C T G A A T A G A G A T G A G C A C T A C A C T A T T G A T G A A C C Majority									
3160		3170		3180		3190		3200	
148	A	T	T	G	G	T	T	C	G
917	A	T	T	G	G	T	T	C	G
149	A	T	T	G	G	T	T	C	G
149	A	T	T	G	G	T	T	C	G
151	A	T	T	G	G	T	T	C	G
150	A	T	T	G	G	T	T	C	G
A A C A G T T G A T A A C C A A A A T A C A T T A A A A A T T A C G T T T A A A C C A G A G A A A T Majority									
3210		3220		3230		3240		3250	
198	A	A	C	A	G	T	T	G	A
967	A	A	C	A	G	T	T	G	A
199	A	A	C	A	G	T	T	G	A
199	A	A	C	A	G	T	T	G	A
201	A	A	C	A	G	T	T	G	A
200	A	A	C	A	G	T	T	G	A
T T A A A G A A A T T G C T G A G C T A C T T A A A G G A A T G A C C C T T G T T A A A A A T C A A Majority									
3260		3270		3280		3290		3300	
248	T	T	A	A	A	G	A	A	T
217	T	T	A	A	A	G	A	A	T
249	T	T	A	A	A	G	A	A	T
249	T	T	A	A	A	G	A	A	T
251	T	T	A	A	A	G	A	A	T
250	T	T	A	A	A	G	A	A	T
G A T G C T C T T G A T A A A G C T A C T G C A A A T A C A G A T G A T G C G G C A T T T T T G G A Majority									
3310		3320		3330		3340		3350	
298	G	A	T	G	C	T	C	T	T
267	G	A	T	G	C	T	C	T	T
299	G	A	T	G	C	T	C	T	T
299	G	A	T	G	C	T	C	T	T
301	G	A	T	G	C	T	C	T	T
300	G	A	T	G	C	T	C	T	T
A A T T C C A G T T G C A T C A A C T A T T A A T G A A A A A G C A G T T T T A G G A A A A G C A A Majority									
3360		3370		3380		3390		3400	
48	A	A	T	T	C	C	A	G	T
17	A	A	T	T	C	C	A	G	T
49	A	A	T	T	C	C	A	G	T
49	A	A	T	T	C	C	A	G	T
51	A	A	T	T	C	C	A	G	T
50	A	A	T	T	C	C	A	G	T
T T G A A A A T A C T T T T T G A A C T T C A A T A T G A C C A T A C T C C T G A T A A A G C T G A C Majority									
3410		3420		3430		3440		3450	
98	T	T	G	A	A	A	T	A	C
67	T	T	G	A	A	A	T	A	C
99	T	T	G	A	A	A	T	A	C
99	T	T	G	A	A	A	T	A	C
01	T	T	G	A	A	A	T	A	C
00	T	T	G	A	A	A	T	A	C
A A T C C A A A A C C A T C T A A T C C T C C A A G A A A A C C A G A A G T T C A T A C T G G T G G Majority									
3460		3470		3480		3490		3500	
48	A	A	T	C	C	A	A	A	A
17	A	A	T	C	C	A	A	A	A
49	A	A	T	C	C	A	A	A	A
49	A	A	T	C	C	A	A	A	A
51	A	A	T	C	C	A	A	A	A
50	A	A	T	C	C	A	A	A	A

FIGURE 18 I

G A A A C G A T T T G T A A A G A A A G A C T C A A C A G A A A C A C A A A C A C T A G G T G G T C Majority									
3510		3520		3530		3540		3550	
1498	G A A A C G A T T T G T A A A G A A A G A C T C A A C A G A A A C A C A A A C A C T A G G T G G T C 2603_all.seq								
1267	G A A A C G A T T T G T A A A G A A A G A C T C A A C A G A A A C A C A A A C A C T A G G T G G T C 18rs21_all.seq								
1499	G A A A C G A T T T G T A A A G A A A G A C T C A A C A G A A A C A C A A A C A C T A G G T G G T C coh1_all.seq								
1499	G A A A C G A T T T G T A A A G A A A G A C T C A A C A G A A A C A C A A A C A C T A G G T G G T C cjb111_all.seq								
1501	G A A A C G A T T T G T A A A G A A A G A C T C A A C A G A A A C A C A A A C A C T A G G T G G T C nem316_all.seq								
1500	G A A A C G A T T T G T A A A G A A A G A C T C A A C A G A A A C A C A A A C A C T A G G T G G T C a909_all.seq								
C T G A G T T T G A T T T G T T G G C T T C T G A T G G G A C A G C A G T A A A A T G G A C A G A T Majority									
3560		3570		3580		3590		3600	
1548	C T G A G T T T G A T T T G T T G G C T T C T G A T G G G A C A G C A G T A A A A T G G A C A G A T 2603_all.seq								
1317	C T G A G T T T G A T T T G T T G G C T T C T G A T G G G A C A G C A G T A A A A T G G A C A G A T 18rs21_all.seq								
1549	C T G A G T T T G A T T T G T T G G C T T C T G A T G G G A C A G C A G T A A A A T G G A C A G A T coh1_all.seq								
1549	C T G A G T T T G A T T T G T T G G C T T C T G A T G G G A C A G C A G T A A A A T G G A C A G A T cjb111_all.seq								
1551	C T G A G T T T G A T T T G T T G G C T T C T G A T G G G A C A G C A G T A A A A T G G A C A G A T nem316_all.seq								
1550	C T G A G T T T G A T T T G T T G G C T T C T G A T G G G A C A G C A G T A A A A T G G A C A G A T a909_all.seq								
G C T C T T A T T A A A G C G A A T A C T A A T A A A A A C T A T A T T G C T G G A G A A G C T G T Majority									
3610		3620		3630		3640		3650	
598	G C T C T T A T T A A A G C G A A T A C T A A T A A A A A C T A T A T T G C T G G A G A A G C T G T 2603_all.seq								
367	G C T C T T A T T A A A G C G A A T A C T A A T A A A A A C T A T A T T G C T G G A G A A G C T G T 18rs21_all.seq								
599	G C T C T T A T T A A A G C G A A T A C T A A T A A A A A C T A T A T T G C T G G A G A A G C T G T coh1_all.seq								
599	G C T C T T A T T A A A G C G A A T A C T A A T A A A A A C T A T A T T G C T G G A G A A G C T G T cjb111_all.seq								
601	G C T C T T A T T A A A G C G A A T A C T A A T A A A A A C T A T A T T G C T G G A G A A G C T G T nem316_all.seq								
600	G C T C T T A T T A A A G C G A A T A C T A A T A A A A A C T A T A T T G C T G G A G A A G C T G T a909_all.seq								
T A C T G G G C A A C C A A T C A A A T T G A A A T C A C A T A C A G A C G G T A C G T T T G A G A Majority									
3660		3670		3680		3690		3700	
648	T A C T G G G C A A C C A A T C A A A T T G A A A T C A C A T A C A G A C G G T A C G T T T G A G A 2603_all.seq								
417	T A C T G G G C A A C C A A T C A A A T T G A A A T C A C A T A C A G A C G G T A C G T T T G A G A 18rs21_all.seq								
649	T A C T G G G C A A C C A A T C A A A T T G A A A T C A C A T A C A G A C G G T A C G T T T G A G A coh1_all.seq								
649	T A C T G G G C A A C C A A T C A A A T T G A A A T C A C A T A C A G A C G G T A C G T T T G A G A cjb111_all.seq								
651	T A C T G G G C A A C C A A T C A A A T T G A A A T C A C A T A C A G A C G G T A C G T T T G A G A nem316_all.seq								
650	T A C T G G G C A A C C A A T C A A A T T G A A A T C A C A T A C A G A C G G T A C G T T T G A G A a909_all.seq								
T T A A A G G T T T G G C T T A T G C A G T T G A T G C G A A T G C A G A G G G T A C A G C A G T A Majority									
3710		3720		3730		3740		3750	
598	T T A A A G G T T T G G C T T A T G C A G T T G A T G C G A A T G C A G A G G G T A C A G C A G T A 2603_all.seq								
467	T T A A A G G T T T G G C T T A T G C A G T T G A T G C G A A T G C A G A G G G T A C A G C A G T A 18rs21_all.seq								
599	T T A A A G G T T T G G C T T A T G C A G T T G A T G C G A A T G C A G A G G G T A C A G C A G T A coh1_all.seq								
599	T T A A A G G T T T G G C T T A T G C A G T T G A T G C G A A T G C A G A G G G T A C A G C A G T A cjb111_all.seq								
701	T T A A A G G T T T G G C T T A T G C A G T T G A T G C G A A T G C A G A G G G T A C A G C A G T A nem316_all.seq								
700	T T A A A G G T T T G G C T T A T G C A G T T G A T G C G A A T G C A G A G G G T A C A G C A G T A a909_all.seq								
A C T T A C A A A T T A A A A G A A A C A A A A G C A C C A G A A G G T T A T G T A A T C C C T G A Majority									
3760		3770		3780		3790		3800	
748	A C T T A C A A A T T A A A A G A A A C A A A A G C A C C A G A A G G T T A T G T A A T C C C T G A 2603_all.seq								
517	A C T T A C A A A T T A A A A G A A A C A A A A G C A C C A G A A G G T T A T G T A A T C C C T G A 18rs21_all.seq								
749	A C T T A C A A A T T A A A A G A A A C A A A A G C A C C A G A A G G T T A T G T A A T C C C T G A coh1_all.seq								
749	A C T T A C A A A T T A A A A G A A A C A A A A G C A C C A G A A G G T T A T G T A A T C C C T G A cjb111_all.seq								
751	A C T T A C A A A T T A A A A G A A A C A A A A G C A C C A G A A G G T T A T G T A A T C C C T G A nem316_all.seq								
750	A C T T A C A A A T T A A A A G A A A C A A A A G C A C C A G A A G G T T A T G T A A T C C C T G A a909_all.seq								
T A A A G A A A T C G A G T T T A C A G T A T C A C A A A C A T C T T A T A A T A C A A A A C C A A Majority									
3810		3820		3830		3840		3850	
798	T A A A G A A A T C G A G T T T A C A G T A T C A C A A A C A T C T T A T A A T A C A A A A C C A A 2603_all.seq								
567	T A A A G A A A T C G A G T T T A C A G T A T C A C A A A C A T C T T A T A A T A C A A A A C C A A 18rs21_all.seq								
799	T A A A G A A A T C G A G T T T A C A G T A T C A C A A A C A T C T T A T A A T A C A A A A C C A A coh1_all.seq								
799	T A A A G A A A T C G A G T T T A C A G T A T C A C A A A C A T C T T A T A A T A C A A A A C C A A cjb111_all.seq								
801	T A A A G A A A T C G A G T T T A C A G T A T C A C A A A C A T C T T A T A A T A C A A A A C C A A nem316_all.seq								
800	T A A A G A A A T C G A G T T T A C A G T A T C A C A A A C A T C T T A T A A T A C A A A A C C A A a909_all.seq								

FIGURE 18 J



**FIGURE 18 K**

T C A G T T G A C G A T T G T T C A T C T T G A A G C A A G G G A T A T T G A T C G T C C A A A T C Majority									
4210		4220		4230		4240		4250	
4198	T	C	A	G	T	T	G	A	C
3967	T	C	A	G	T	T	G	A	C
4199	T	C	A	G	T	T	G	A	C
4199	T	C	A	G	T	T	G	A	C
4201	T	C	A	G	T	T	G	A	C
4200	T	C	A	G	T	T	G	A	C
C A C A G T T G G A G A T T G C C C C T A A A G A A G G G A C T C C A A T T G A A G G A G T A C T C Majority									
4260		4270		4280		4290		4300	
4248	C	A	C	A	G	T	T	G	A
4017	C	A	C	A	G	T	T	G	A
4249	C	A	C	A	G	T	T	G	A
4249	C	A	C	A	G	T	T	G	A
4251	C	A	C	A	G	T	T	G	A
4250	C	A	C	A	G	T	T	G	A
T A T C A G T T G T A C C A A T T A A A A T C A A C T G A A G A T G C C G A T T T G T T G G C A C A Majority									
4310		4320		4330		4340		4350	
4298	T	A	T	C	A	G	T	T	G
4067	T	A	T	C	A	G	T	T	G
4299	T	A	T	C	A	G	T	T	G
4299	T	A	T	C	A	G	T	T	G
4301	T	A	T	C	A	G	T	T	G
4300	T	A	T	C	A	G	T	T	G
T T G G A A T T C C C T A A C T A T C A C A G A A T T G A A A A A A C A G G C G C A G C A G G T T T Majority									
4360		4370		4380		4390		4400	
4348	T	T	G	G	A	A	T	T	C
4117	T	T	G	G	A	A	T	T	C
4349	T	T	G	G	A	A	T	T	C
4349	T	T	G	G	A	A	T	T	C
4351	T	T	G	G	A	A	T	T	C
4350	T	T	G	G	A	A	T	T	C
T T G A A G C C A C T A C T A A T C A A C A A G G A A A G G C T A C A T T T A A C C A A C T A C C A Majority									
4410		4420		4430		4440		4450	
4398	T	T	G	A	A	G	C	C	A
4167	T	T	G	A	A	G	C	C	A
4399	T	T	G	A	A	G	C	C	A
4399	T	T	G	A	A	G	C	C	A
4401	T	T	G	A	A	G	C	C	A
4400	T	T	G	A	A	G	C	C	A
G A T G G A A T T T A T T A T G G T C T G G C G G T T A A A G C C G G T G A A A A A A A T C G T A A Majority									
4460		4470		4480		4490		4500	
448	G	A	T	G	G	A	A	T	T
217	G	A	T	G	G	A	A	T	T
449	G	A	T	G	G	A	A	T	T
449	G	A	T	G	G	A	A	T	T
451	G	A	T	G	G	A	A	T	T
450	G	A	T	G	G	A	A	T	T
T G T C T C A G C T T T C T T G G T T G A C T T G T C T G A G G A T A A A G T G A T T T A T C C T A Majority									
4510		4520		4530		4540		4550	
498	T	G	T	C	T	C	A	G	C
267	T	G	T	C	T	C	A	G	C
499	T	G	T	C	T	C	A	G	C
499	T	G	T	C	T	C	A	G	C
501	T	G	T	C	T	C	A	G	C
500	T	G	T	C	T	C	A	G	C

FIGURE 18 L



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A A A T C A T C T G G T C C A C A G G T G A G T T G G A C T T G C T T A A A G T T G G T G T G G A T Majority									
4560		4570		4580		4590		4600	
4548	A A A T C A T C T G G T C C A C A G G T G A G T T G G A C T T G C T T A A A G T T G G T G T G G A T 2603_ail.seq								
4317	A A A T C A T C T G G T C C A C A G G T G A G T T G G A C T T G C T T A A A G T T G G T G T G G A T 18rs21_ail.seq								
4549	A A A T C A T C T G G T C C A C A G G T G A G T T G G A C T T G C T T A A A G T T G G T G T G G A T coh1_ail.seq								
4549	A A A T C A T C T G G T C C A C A G G T G A G T T G G A C T T G C T T A A A G T T G G T G T G G A T cjb111_ail.seq								
4551	A A A T C A T C T G G T C C A C A G G T G A G T T G G A C T T G C T T A A A G T T G G T G T G G A T nem316_ail.seq								
4550	A A A T C A T C T G G T C C A C A G G T G A G T T G G A C T T G C T T A A A G T T G G T G T G G A T a909_ail.seq								
G G T G A T A C C A A A A A A C C A C T A G C A G G C G T T G T C T T T G A A C T T T A T G A A A A Majority									
4610		4620		4630		4640		4650	
1598	G G T G A T A C C A A A A A A C C A C T A G C A G G C G T T G T C T T T G A A C T T T A T G A A A A 2603_ail.seq								
1367	G G T G A T A C C A A A A A A C C A C T A G C A G G C G T T G T C T T T G A A C T T T A T G A A A A 18rs21_ail.seq								
1599	G G T G A T A C C A A A A A A C C A C T A G C A G G C G T T G T C T T T G A A C T T T A T G A A A A coh1_ail.seq								
1599	G G T G A T A C C A A A A A A C C A C T A G C A G G C G T T G T C T T T G A A C T T T A T G A A A A cjb111_ail.seq								
1601	G G T G A T A C C A A A A A A C C A C T A G C A G G C G T T G T C T T T G A A C T T T A T G A A A A nem316_ail.seq								
1600	G G T G A T A C C A A A A A A C C A C T A G C A G G C G T T G T C T T T G A A C T T T A T G A A A A a909_ail.seq								
G A A T G G T A G G A C T C C T A T T C G T G T G A A A A A T G G G G T G C A T T C T C A A G A T A Majority									
4660		4670		4680		4690		4700	
1648	G A A T G G T A G G A C T C C T A T T C G T G T G A A A A A T G G G G T G C A T T C T C A A G A T A 2603_ail.seq								
1417	G A A T G G T A G G A C T C C T A T T C G T G T G A A A A A T G G G G T G C A T T C T C A A G A T A 18rs21_ail.seq								
1649	G A A T G G T A G G A C T C C T A T T C G T G T G A A A A A T G G G G T G C A T T C T C A A G A T A coh1_ail.seq								
1649	G A A T G G T A G G A C T C C T A T T C G T G T G A A A A A T G G G G T G C A T T C T C A A G A T A cjb111_ail.seq								
1651	G A A T G G T A G G A C T C C T A T T C G T G T G A A A A A T G G G G T G C A T T C T C A A G A T A nem316_ail.seq								
1650	G A A T G G T A G G A C T C C T A T T C G T G T G A A A A A T G G G G T G C A T T C T C A A G A T A a909_ail.seq								
T T G A C G C T G C A A A A C A T T T A G A A A C A G A T T C A T C A G G G C A T A T C A G A A T T Majority									
4710		4720		4730		4740		4750	
1698	T T G A C G C T G C A A A A C A T T T A G A A A C A G A T T C A T C A G G G C A T A T C A G A A T T 2603_ail.seq								
1467	T T G A C G C T G C A A A A C A T T T A G A A A C A G A T T C A T C A G G G C A T A T C A G A A T T 18rs21_ail.seq								
1699	T T G A C G C T G C A A A A C A T T T A G A A A C A G A T T C A T C A G G G C A T A T C A G A A T T coh1_ail.seq								
1699	T T G A C G C T G C A A A A C A T T T A G A A A C A G A T T C A T C A G G G C A T A T C A G A A T T cjb111_ail.seq								
1701	T T G A C G C T G C A A A A C A T T T A G A A A C A G A T T C A T C A G G G C A T A T C A G A A T T nem316_ail.seq								
1700	T T G A C G C T G C A A A A C A T T T A G A A A C A G A T T C A T C A G G G C A T A T C A G A A T T a909_ail.seq								
T C C G G G C T C A T C C A T G G G G A C T A T G T C T T A A A A G A A A T C G A G A C A C A G T C Majority									
4760		4770		4780		4790		4800	
1748	T C C G G G C T C A T C C A T G G G G A C T A T G T C T T A A A A G A A A T C G A G A C A C A G T C 2603_ail.seq								
1517	T C C G G G C T C A T C C A T G G G G A C T A T G T C T T A A A A G A A A T C G A G A C A C A G T C 18rs21_ail.seq								
1749	T C C G G G C T C A T C C A T G G G G A C T A T G T C T T A A A A G A A A T C G A G A C A C A G T C coh1_ail.seq								
1749	T C C G G G C T C A T C C A T G G G G A C T A T G T C T T A A A A G A A A T C G A G A C A C A G T C cjb111_ail.seq								
1751	T C C G G G C T C A T C C A T G G G G A C T A T G T C T T A A A A G A A A T C G A G A C A C A G T C nem316_ail.seq								
1750	T C C G G G C T C A T C C A T G G G G A C T A T G T C T T A A A A G A A A T C G A G A C A C A G T C a909_ail.seq								
A G G A T A T C A G A T C G G A C A G G C A G A G A C T G C T G T G A C T A T T G A A A A A T C A A Majority									
4810		4820		4830		4840		4850	
1798	A G G A T A T C A G A T C G G A C A G G C A G A G A C T G C T G T G A C T A T T G A A A A A T C A A 2603_ail.seq								
1567	A G G A T A T C A G A T C G G A C A G G C A G A G A C T G C T G T G A C T A T T G A A A A A T C A A 18rs21_ail.seq								
1799	A G G A T A T C A G A T C G G A C A G G C A G A G A C T G C T G T G A C T A T T G A A A A A T C A A coh1_ail.seq								
1799	A G G A T A T C A G A T C G G A C A G G C A G A G A C T G C T G T G A C T A T T G A A A A A T C A A cjb111_ail.seq								
1801	A G G A T A T C A G A T C G G A C A G G C A G A G A C T G C T G T G A C T A T T G A A A A A T C A A nem316_ail.seq								
1800	A G G A T A T C A G A T C G G A C A G G C A G A G A C T G C T G T G A C T A T T G A A A A A T C A A a909_ail.seq								
A A A C A G T A A C A G T A A C G A T T G A A A A T A A A A A A G T T C C G A C A C C T A A A G T C Majority									
4860		4870		4880		4890		4900	
1848	A A A C A G T A A C A G T A A C G A T T G A A A A T A A A A A A G T T C C G A C A C C T A A A G T C 2603_ail.seq								
1617	A A A C A G T A A C A G T A A C G A T T G A A A A T A A A A A A G T T C C G A C A C C T A A A G T C 18rs21_ail.seq								
1849	A A A C A G T A A C A G T A A C G A T T G A A A A T A A A A A A G T T C C G A C A C C T A A A G T C coh1_ail.seq								
1849	A A A C A G T A A C A G T A A C G A T T G A A A A T A A A A A A G T T C C G A C A C C T A A A G T C cjb111_ail.seq								
1851	A A A C A G T A A C A G T A A C G A T T G A A A A T A A A A A A G T T C C G A C A C C T A A A G T C nem316_ail.seq								
1850	A A A C A G T A A C A G T A A C G A T T G A A A A T A A A A A A G T T C C G A C A C C T A A A G T C a909_ail.seq								

FIGURE 18 M

C C A T C T C G A G G A G G T C T T A T T C C C A A A A C A G G T G A G C A A C A G G C A A T G G C Majority									
4910		4920		4930		4940		4950	
4898	C C A T C T C G A G G A G G T C T T A T T C C C A A A A C A G G T G A G C A A C A G G C A A T G G C 2603_all.seq								
4667	C C A T C T C G A G G A G G T C T T A T T C C C A A A A C A G G T G A G C A A C A G G C A A T G G C 18rs21_all.seq								
4899	C C A T C T C G A G G A G G T C T T A T T C C C A A A A C A G G T G A G C A A C A G G C A A T G G C coh1_all.seq								
4899	C C A T C T C G A G G A G G T C T T A T T C C C A A A A C A G G T G A G C A A C A G G C A A T G G C cjb111_all.seq								
4901	C C A T C T C G A G G A G G T C T T A T T C C C A A A A C A G G T G A G C A A C A G G C A A T G G C nem316_all.seq								
4900	C C A T C T C G A G G A G G T C T T A T T C C C A A A A C A G G T G A G C A A C A G G C A A T G G C a909_all.seq								
A C T T G T A A T T A T T G G T G G T A T T T T A A T T G C T T T A G C C T T A C G A T T A C T A T Majority									
4960		4970		4980		4990		5000	
4948	A C T T G T A A T T A T T G G T G G T A T T T T A A T T G C T T T A G C C T T A C G A T T A C T A T 2603_all.seq								
4717	A C T T G T A A T T A T T G G T G G T A T T T T A A T T G C T T T A G C C T T A C G A T T A C T A T 18rs21_all.seq								
4949	A C T T G T A A T T A T T G G T G G T A T T T T A A T T G C T T T A G C C T T A C G A T T A C T A T coh1_all.seq								
4949	A C T T G T A A T T A T T G G T G G T A T T T T A A T T G C T T T A G C C T T A C G A T T A C T A T cjb111_all.seq								
4951	A C T T G T A A T T A T T G G T G G T A T T T T A A T T G C T T T A G C C T T A C G A T T A C T A T nem316_all.seq								
4950	A C T T G T A A T T A T T G G T G G T A T T T T A A T T G C T T T A G C C T T A C G A T T A C T A T a909_all.seq								
C A A A A C A T C G G A A A C A T C A A A A T A A G C A T T A G C A T G G G A C A A A A A T C A A A Majority									
5010		5020		5030		5040		5050	
4998	C A A A A C A T C G G A A A C A T C A A A A T A A G G A T T A G C A T G G G A C A A A A A T C A A A 2603_all.seq								
4767	C A A A A C A T C G G A A A C A T C A A A A T A A G G A T T A G C A T G G G A C A A A A A T C A A A 18rs21_all.seq								
4999	C A A A A C A T C G G A A A C A T C A A A A T A A G G A T T A G C A T G G G A C A A A A A T C A A A coh1_all.seq								
4999	C A A A A C A T C G G A A A C A T C A A A A T A A G G A T T A G C A T G G G A C A A A A A T C A A A cjb111_all.seq								
5001	C A A A A C A T C G G A A A C A T C A A A A T A A G G A T T A G C A T G G G A C A A A A A T C A A A nem316_all.seq								
5000	C A A A A C A T C G G A A A C A T C A A A A T A A G G A T T A G C A T G G G A C A A A A A T C A A A a909_all.seq								
A A T A T C T C T A G C T A C G A A T A T T C G T A T A T G G A T T T T T C G T T T A A T T T T C T Majority									
5060		5070		5080		5090		5100	
5048	A A T A T C T C T A G C T A C G A A T A T T C G T A T A T G G A T T T T T C G T T T A A T T T T C T 2603_all.seq								
1817	A A T A T C T C T A G C T A C G A A T A T T C G T A T A T G G A T T T T T C G T T T A A T T T T C T 18rs21_all.seq								
5049	A A T A T C T C T A G C T A C G A A T A T T C G T A T A T G G A T T T T T C G T T T A A T T T T C T coh1_all.seq								
5049	A A T A T C T C T A G C T A C G A A T A T T C G T A T A T G G A T T T T T C G T T T A A T T T T C T cjb111_all.seq								
5051	A A T A T C T C T A G C T A C G A A T A T T C G T A T A T G G A T T T T T C G T T T A A T T T T C T nem316_all.seq								
5050	A A T A T C T C T A G C T A C G A A T A T T C G T A T A T G G A T T T T T C G T T T A A T T T T C T a909_all.seq								
T A G C G G G T T T C C T T G T T T T G G C A T T T C C C A T C G T T A G T C A G G T C A T G T A C Majority									
5110		5120		5130		5140		5150	
5098	T A G C G G G T T T C C T T G T T T T G G C A T T T C C C A T C G T T A G T C A G G T C A T G T A C 2603_all.seq								
1867	T A G C G G G T T T C C T T G T T T T G G C A T T T C C C A T C G T T A G T C A G G T C A T G T A C 18rs21_all.seq								
5099	T A G C G G G T T T C C T T G T T T T G G C A T T T C C C A T C G T T A G T C A G G T C A T G T A C coh1_all.seq								
5099	T A G C G G G T T T C C T T G T T T T G G C A T T T C C C A T C G T T A G T C A G G T C A T G T A C cjb111_all.seq								
5101	T A G C G G G T T T C C T T G T T T T G G C A T T T C C C A T C G T T A G T C A G G T C A T G T A C nem316_all.seq								
5100	T A G C G G G T T T C C T T G T T T T G G C A T T T C C C A T C G T T A G T C A G G T C A T G T A C a909_all.seq								
T T T C A A G C C T C T C A C G C C A A T A T T A A T G C T T T T A A A G A A G C T G T T A C C A A Majority									
5160		5170		5180		5190		5200	
5148	T T T C A A G C C T C T C A C G C C A A T A T T A A T G C T T T T A A A G A A G C T G T T A C C A A 2603_all.seq								
917	T T T C A A G C C T C T C A C G C C A A T A T T A A T G C T T T T A A A G A A G C T G T T A C C A A 18rs21_all.seq								
5149	T T T C A A G C C T C T C A C G C C A A T A T T A A T G C T T T T A A A G A A G C T G T T A C C A A coh1_all.seq								
5149	T T T C A A G C C T C T C A C G C C A A T A T T A A T G C T T T T A A A G A A G C T G T T A C C A A cjb111_all.seq								
5151	T T T C A A G C C T C T C A C G C C A A T A T T A A T G C T T T T A A A G A A G C T G T T A C C A A nem316_all.seq								
5150	T T T C A A G C C T C T C A C G C C A A T A T T A A T G C T T T T A A A G A A G C T G T T A C C A A a909_all.seq								
G A T T G A C C G G G T G G A G A T T A A T C G G C G T T T A G A A C T T G C T T A T G C T T A T A Majority									
5210		5220		5230		5240		5250	
198	G A T T G A C C G G G T G G A G A T T A A T C G G C G T T T A G A A C T T G C T T A T G C T T A T A 2603_all.seq								
967	G A T T G A C C G G G T G G A G A T T A A T C G G C G T T T A G A A C T T G C T T A T G C T T A T A 18rs21_all.seq								
199	G A T T G A C C G G G T G G A G A T T A A T C G G C G T T T A G A A C T T G C T T A T G C T T A T A coh1_all.seq								
199	G A T T G A C C G G G T G G A G A T T A A T C G G C G T T T A G A A C T T G C T T A T G C T T A T A cjb111_all.seq								
201	G A T T G A C C G G G T G G A G A T T A A T C G G C G T T T A G A A C T T G C T T A T G C T T A T A nem316_all.seq								
200	G A T T G A C C G G G T G G A G A T T A A T C G G C G T T T A G A A C T T G C T T A T G C T T A T A a909_all.seq								

FIGURE 18 N



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A C G C C A G T A T A G C A G G T G C C A A A A C T A A T G G C G A A T A T C C A G C G C T T A A A Majority																														
5260						5270						5280						5290						5300						
5248	A C G C C A G T A T A G C A G G T G C C A A A A C T A A T G G C G A A T A T C C A G C G C T T A A A																													2603_all.seq
5017	A C G C C A G T A T A G C A G G T G C C A A A A C T A A T G G C G A A T A T C C A G C G C T T A A A																													18rs21_all.seq
5249	A C G C C A G T A T A G C A G G T G C C A A A A C T A A T G G C G A A T A T C C A G C G C T T A A A																													cohl1_all.seq
5249	A C G C C A G T A T A G C A G G T G C C A A A A C T A A T G G C G A A T A T C C A G C G C T T A A A																													cjb111_all.seq
5251	A C G C C A G T A T A G C A G G T G C C A A A A C T A A T G G C G A A T A T C C A G C G C T T A A A																													nen316_all.seq
5250	A C G C C A G T A T A G C A G G T G C C A A A A C T A A T G G C G A A T A T C C A G C G C T T A A A																													a909_all.seq
G A C C C C T A C T C T G C T G A A C A A A A G C A G G C A G G G G T C G T T G A G T A C G C C C G Majority																														
5310						5320						5330						5340						5350						
5298	G A C C C C T A C T C T G C T G A A C A A A A G C A G G C A G G G G T C G T T G A G T A C G C C C G																													2603_all.seq
5067	G A C C C C T A C T C T G C T G A A C A A A A G C A G G C A G G G G T C G T T G A G T A C G C C C G																													18rs21_all.seq
5299	G A C C C C T A C T C T G C T G A A C A A A A G C A G G C A G G G G T C G T T G A G T A C G C C C G																													cohl1_all.seq
5299	G A C C C C T A C T C T G C T G A A C A A A A G C A G G C A G G G G T C G T T G A G T A C G C C C G																													cjb111_all.seq
5301	G A C C C C T A C T C T G C T G A A C A A A A G C A G G C A G G G G T C G T T G A G T A C G C C C G																													nen316_all.seq
5300	G A C C C C T A C T C T G C T G A A C A A A A G C A G G C A G G G G T C G T T G A G T A C G C C C G																													a909_all.seq
C A T G C T T G A A G T C A A A G A A C A A A T A G G T C A T G T G A T T A T T C C A A G A A T T A Majority																														
5360						5370						5380						5390						5400						
5348	C A T G C T T G A A G T C A A A G A A C A A A T A G G T C A T G T G A T T A T T C C A A G A A T T A																													2603_all.seq
5117	C A T G C T T G A A G T C A A A G A A C A A A T A G G T C A T G T G A T T A T T C C A A G A A T T A																													18rs21_all.seq
5349	C A T G C T T G A A G T C A A A G A A C A A A T A G G T C A T G T G A T T A T T C C A A G A A T T A																													cohl1_all.seq
5349	C A T G C T T G A A G T C A A A G A A C A A A T A G G T C A T G T G A T T A T T C C A A G A A T T A																													cjb111_all.seq
5351	C A T G C T T G A A G T C A A A G A A C A A A T A G G T C A T G T G A T T A T T C C A A G A A T T A																													nen316_all.seq
5350	C A T G C T T G A A G T C A A A G A A C A A A T A G G T C A T G T G A T T A T T C C A A G A A T T A																													a909_all.seq
A T C A G G A T A T C C C T A T T T A C G C T G G C T C T G C T G A A G A A A A T C T T C A G A G G Majority																														
5410						5420						5430						5440						5450						
5398	A T C A G G A T A T C C C T A T T T A C G C T G G C T C T G C T G A A G A A A A T C T T C A G A G G																													2603_all.seq
5167	A T C A G G A T A T C C C T A T T T A C G C T G G C T C T G C T G A A G A A A A T C T T C A G A G G																													18rs21_all.seq
5399	A T C A G G A T A T C C C T A T T T A C G C T G G C T C T G C T G A A G A A A A T C T T C A G A G G																													cohl1_all.seq
5399	A T C A G G A T A T C C C T A T T T A C G C T G G C T C T G C T G A A G A A A A T C T T C A G A G G																													cjb111_all.seq
5401	A T C A G G A T A T C C C T A T T T A C G C T G G C T C T G C T G A A G A A A A T C T T C A G A G G																													nen316_all.seq
5400	A T C A G G A T A T C C C T A T T T A C G C T G G C T C T G C T G A A G A A A A T C T T C A G A G G																													a909_all.seq
G G C G T T G G A C A T T T A G A G G G G A C C A G T C T T C C A G T C G G T G G T G A G T C A A C Majority																														
5460						5470						5480						5490						5500						
448	G G C G T T G G A C A T T T A G A G G G G A C C A G T C T T C C A G T C G G T G G T G A G T C A A C																													2603_all.seq
217	G G C G T T G G A C A T T T A G A G G G G A C C A G T C T T C C A G T C G G T G G T G A G T C A A C																													18rs21_all.seq
449	G G C G T T G G A C A T T T A G A G G G G A C C A G T C T T C C A G T C G G T G G T G A G T C A A C																													cohl1_all.seq
449	G G C G T T G G A C A T T T A G A G G G G A C C A G T C T T C C A G T C G G T G G T G A G T C A A C																													cjb111_all.seq
451	G G C G T T G G A C A T T T A G A G G G G A C C A G T C T T C C A G T C G G T G G T G A G T C A A C																													nen316_all.seq
450	G G C G T T G G A C A T T T A G A G G G G A C C A G T C T T C C A G T C G G T G G T G A G T C A A C																													a909_all.seq
T C A T G C C G T T C T A A C T G C C C A T C G A G G G C T A C C A A C G G C C A A G C T A T T T A Majority																														
5510						5520						5530						5540						5550						
498	T C A T G C C G T T C T A A C T G C C C A T C G A G G G C T A C C A A C G G G C A A G C T A T T T A																													2603_all.seq
267	T C A T G C C G T T C T A A C T G C C C A T C G A G G G C T A C C A A C G G C C A A G C T A T T T A																													18rs21_all.seq
499	T C A T G C C G T T C T A A C T G C C C A T C G A G G G C T A C C A A C G G G C A A G C T A T T T A																													cohl1_all.seq
499	T C A T G C C G T T C T A A C T G C C C A T C G A G G G C T A C C A A C G G C C A A G C T A T T T A																													cjb111_all.seq
501	T C A T G C C G T T C T A A C T G C C C A T C G A G G G C T A C C A A C G G C C A A G C T A T T T A																													nen316_all.seq
500	T C A T G C C G T T C T A A C T G C C C A T C G A G G G C T A C C A A C G G C C A A G C T A T T T A																													a909_all.seq
C C A A T T T A G A C A A G G T A A C A G T A G G T G A C C G T T T T T A C A T T G A A C A C A T C Majority																														
5560						5570						5580						5590						5600						
548	C C A A T T T A G A C A A G G T A A C A G T A G G T G A C C G T T T T T A C A T T G A A C A C A T C																													2603_all.seq
517	C C A A T T T A G A C A A G G T A A C A G T A G G T G A C C G T T T T T A C A T T G A A C A C A T C																													18rs21_all.seq
549	C C A A T T T A G A C A A G G T A A C A G T A G G T G A C C G T T T T T A C A T T G A A C A C A T C																													cohl1_all.seq
549	C C A A T T T A G A C A A G G T A A C A G T A G G T G A C C G T T T T T A C A T T G A A C A C A T C																													cjb111_all.seq
551	C C A A T T T A G A C A A G G T A A C A G T A G G T G A C C G T T T T T A C A T T G A A C A C A T C																													nen316_all.seq
550	C C A A T T T A G A C A A G G T A A C A G T A G G T G A C C G T T T T T A C A T T G A A C A C A T C																													a909_all.seq

**FIGURE 18 P**



A T A A A A T C A G A A C C C T C A T T T T T G T G A T G G G A A G T C T G A T T C T C T T A T T T Majority									
5960		5970		5980		5990		6000	
5948	A T A A A A T C A G A A C C C T C A T T T T T G T G A T G G G A A G T C T G A T T C T C T T A T T T 2603_all.seq								
5717	A T A A A A T C A G A A C C C T C A T T T T T G T G A T G G G A A G T C T G A T T C T C T T A T T T 18rs21_all.seq								
5949	A T A A A A T C A G A A C C C T C A T T T T T G T G A T G G G A A G T C T G A T T C T C T T A T T T coh1_all.seq								
5949	A T A A A A T C A G A A C C C T C A T T T T T G T G A T G G G A A G T C T G A T T C T C T T A T T T cjb111_all.seq								
5951	A T A A A A T C A G A A C C C T C A T T T T T G T G A T G G G A A G T C T G A T T C T C T T A T T T nem316_all.seq								
5950	A T A A A A T C A G A A C C C T C A T T T T T G T G A T G G G A A G T C T G A T T C T C T T A T T T a909_all.seq								
C C G A T T G T G A G C C A G G T A A G T T A C T A C C T T G C T T C G C A T C A A A A T A T T A A Majority									
6010		6020		6030		6040		6050	
5998	C C G A T T G T G A G C C A G G T A A G T T A C T A C C T T G C T T C G C A T C A A A A T A T T A A 2603_all.seq								
5767	C C G A T T G T G A G C C A G G T A A G T T A C T A C C T T G C T T C G C A T C A A A A T A T T A A 18rs21_all.seq								
5999	C C G A T T G T G A G C C A G G T A A G T T A C T A C C T T G C T T C G C A T C A A A A T A T T A A coh1_all.seq								
5999	C C G A T T G T G A G C C A G G T A A G T T A C T A C C T T G C T T C G C A T C A A A A T A T T A A cjb111_all.seq								
5001	C C G A T T G T G A G C C A G G T A A G T T A C T A C C T T G C T T C G C A T C A A A A T A T T A A nem316_all.seq								
5000	C C G A T T G T G A G C C A G G T A A G T T A C T A C C T T G C T T C G C A T C A A A A T A T T A A a909_all.seq								
T C A A T T T A A G C G G G A A G T C G C T A A G A T T G A T A C T A A T A C G G T T G A A C G A C Majority									
6060		6070		6080		6090		6100	
5048	T C A A T T T A A G C G G G A A G T C G C T A A G A T T G A T A C T A A T A C G G T T G A A C G A C 2603_all.seq								
5817	T C A A T T T A A G C G G G A A G T C G C T A A G A T T G A T A C T A A T A C G G T T G A A C G A C 18rs21_all.seq								
5049	T C A A T T T A A G C G G G A A G T C G C T A A G A T T G A T A C T A A T A C G G T T G A A C G A C coh1_all.seq								
5049	T C A A T T T A A G C G G G A A G T C G C T A A G A T T G A T A C T A A T A C G G T T G A A C G A C cjb111_all.seq								
5051	T C A A T T T A A G C G G G A A G T C G C T A A G A T T G A T A C T A A T A C G G T T G A A C G A C nem316_all.seq								
5050	T C A A T T T A A G C G G G A A G T C G C T A A G A T T G A T A C T A A T A C G G T T G A A C G A C a909_all.seq								
G C A T C G C T T T A G C T A A T G C T T A C A A T G A C A C G T T A T C A A G G A A T C C C T T G Majority									
6110		6120		6130		6140		6150	
5098	G C A T C G C T T T A G C T A A T G C T T A C A A T G A G A C G T T A T C A A G G A A T C C C T T G 2603_all.seq								
5867	G C A T C G C T T T A G C T A A T G C T T A C A A T G A G A C G T T A T C A A G G A A T C C C T T G 18rs21_all.seq								
5099	G C A T C G C T T T A G C T A A T G C T T A C A A T G A G A C G T T A T C A A G G A A T C C C T T G coh1_all.seq								
5099	G C A T C G C T T T A G C T A A T G C T T A C A A T G A G A C G T T A T C A A G G A A T C C C T T G cjb111_all.seq								
5101	G C A T C G C T T T A G C T A A T G C T T A C A A T G A G A C G T T A T C A A G G A A T C C C T T G nem316_all.seq								
5100	G C A T C G C T T T A G C T A A T G C T T A C A A T G A G A C G T T A T C A A G G A A T C C C T T G a909_all.seq								
C T T A T A G A C C C T T T T A C C A G T A A G C A A A A A G A A G G T T T G A G A G A G T A T G C Majority									
6160		6170		6180		6190		6200	
5148	C T T A T A G A C C C T T T T A C C A G T A A G C A A A A A G A A G G T T T G A G A G A G T A T G C 2603_all.seq								
517	C T T A T A G A C C C T T T T A C C A G T A A G C A A A A A G A A G G T T T G A G A G A G T A T G C 18rs21_all.seq								
5149	C T T A T A G A C C C T T T T A C C A G T A A G C A A A A A G A A G G T T T G A G A G A G T A T G C coh1_all.seq								
5149	C T T A T A G A C C C T T T T A C C A G T A A G C A A A A A G A A G G T T T G A G A G A G T A T G C cjb111_all.seq								
5151	C T T A T A G A C C C T T T T A C C A G T A A G C A A A A A G A A G G T T T G A G A G A G T A T G C nem316_all.seq								
5150	C T T A T A G A C C C T T T T A C C A G T A A G C A A A A A G A A G G T T T G A G A G A G T A T G C a909_all.seq								
T C G T A T G C T T G A A G T T C A T G A G C A A A T A G G T C A T G T G G C A A T C C C A A G T A Majority									
6210		6220		6230		6240		6250	
5198	T C G T A T G C T T G A A G T T C A T G A G C A A A T A G G T C A T G T G G C A A T C C C A A G T A 2603_all.seq								
5967	T C G T A T G C T T G A A G T T C A T G A G C A A A T A G G T C A T G T G G C A A T C C C A A G T A 18rs21_all.seq								
5199	T C G T A T G C T T G A A G T T C A T G A G C A A A T A G G T C A T G T G G C A A T C C C A A G T A coh1_all.seq								
5199	T C G T A T G C T T G A A G T T C A T G A G C A A A T A G G T C A T G T G G C A A T C C C A A G T A cjb111_all.seq								
5201	T C G T A T G C T T G A A G T T C A T G A G C A A A T A G G T C A T G T G G C A A T C C C A A G T A nem316_all.seq								
5200	T C G T A T G C T T G A A G T T C A T G A G C A A A T A G G T C A T G T G G C A A T C C C A A G T A a909_all.seq								
T T G C G G T T G A T A T T C C A A T T T A T G C T G G A A C A T C C G A A A C T G T G C T T C A G Majority									
6260		6270		6280		6290		6300	
5248	T T G C G G T T G A T A T T C C A A T T T A T G C T G G A A C A T C C G A A A C T G T G C T T C A G 2603_all.seq								
517	T T G C G G T T G A T A T T C C A A T T T A T G C T G G A A C A T C C G A A A C T G T G C T T C A G 18rs21_all.seq								
5249	T T G C G G T T G A T A T T C C A A T T T A T G C T G G A A C A T C C G A A A C T G T G C T T C A G coh1_all.seq								
5249	T T G C G G T T G A T A T T C C A A T T T A T G C T G G A A C A T C C G A A A C T G T G C T T C A G cjb111_all.seq								
5251	T T G C G G T T G A T A T T C C A A T T T A T G C T G G A A C A T C C G A A A C T G T G C T T C A G nem316_all.seq								
5250	T T G C G G T T G A T A T T C C A A T T T A T G C T G G A A C A T C C G A A A C T G T G C T T C A G a909_all.seq								

FIGURE 18 Q

A A A G G T A G T G G G C A T T T T G G A G G G A A C C A G T C T T C C A G T G G G A G G T T T G T C Majority									
6310		6320		6330		6340		6350	
6298	A	A	A	G	G	T	A	G	T
6067	A	A	A	G	G	T	A	G	T
6299	A	A	A	G	G	T	A	G	T
6299	A	A	A	G	G	T	A	G	T
6301	A	A	A	G	G	T	A	G	T
6300	A	A	A	G	G	T	A	G	T
A A C C C A T T C A G T A C T A A C T G C C C A C C G T G G C T T G C C A A C A G C T A G G C T A T Majority									
6360		6370		6380		6390		6400	
6348	A	A	C	C	C	A	T	T	C
6117	A	A	C	C	C	A	T	T	C
6349	A	A	C	C	C	A	T	T	C
6349	A	A	C	C	C	A	T	T	C
6351	A	A	C	C	C	A	T	T	C
6350	A	A	C	C	C	A	T	T	C
T T A C C G A C T T A A A T A A A G T T A A A A A A G G C C A G A T T T T C T A T G T G A C G A A C Majority									
6410		6420		6430		6440		6450	
6398	T	T	A	C	C	G	A	C	T
6167	T	T	A	C	C	G	A	C	T
6399	T	T	A	C	C	G	A	C	T
6399	T	T	A	C	C	G	A	C	T
6401	T	T	A	C	C	G	A	C	T
6400	T	T	A	C	C	G	A	C	T
A T C A A G G A A A C A C T T G C C T A C A A A G T C G T G T C T A T C A A A G T T G T G G A T C C Majority									
6460		6470		6480		6490		6500	
6448	A	T	C	A	A	G	G	A	A
6217	A	T	C	A	A	G	G	A	A
6449	A	T	C	A	A	G	G	A	A
6449	A	T	C	A	A	G	G	A	A
6451	A	T	C	A	A	G	G	A	A
6450	A	T	C	A	A	G	G	A	A
A A C A G C T T T A A G T G A G G T T A A G A T T G T C A A T G G T A A G G A T T A T A T A A C C T Majority									
6510		6520		6530		6540		6550	
3498	A	A	C	A	G	C	T	T	T
3267	A	A	C	A	G	C	T	T	T
3499	A	A	C	A	G	C	T	T	T
3499	A	A	C	A	G	C	T	T	T
3501	A	A	C	A	G	C	T	T	T
3500	A	A	C	A	G	C	T	T	T
T G C T G A C T T G C A C A C C T T A C A T G A T C A A T A G T C A T C G T C T C T T G G T A A A A Majority									
6560		6570		6580		6590		6600	
1548	T	G	C	T	G	A	C	T	T
1317	T	G	C	T	G	A	C	T	T
1549	T	G	C	T	G	A	C	T	T
1549	T	G	C	T	G	A	C	T	T
1551	T	G	C	T	G	A	C	T	T
1550	T	G	C	T	G	A	C	T	T
G G A G A G C G T A T T C C T T A T G A T T C T A C C G A G G C G G A A A A G C A C A A A G A A C A Majority									
6610		6620		6630		6640		6650	
1598	G	G	A	G	A	G	C	G	T
1367	G	G	A	G	A	G	C	G	T
1599	G	G	A	G	A	G	C	G	T
1599	G	G	A	G	A	G	C	G	T
1601	G	G	A	G	A	G	C	G	T
1600	G	G	A	G	A	G	C	G	T

FIGURE 18 R



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A A C C G T A C A A G A T T A T C G T T T G T C A C T A G T G T T G A A G A T A C T A C T A G T A T Majority									
6660		6670		6680		6690		6700	
648	A	A	C	C	G	T	A	C	A
647	A	A	C	C	G	T	A	C	A
649	A	A	C	C	G	T	A	C	A
649	A	A	C	C	G	T	A	C	A
651	A	A	C	C	G	T	A	C	A
650	A	A	C	C	G	T	A	C	A
T A T T A A T T G G A C T C T T C A T C G T G A T A A T G A T G A G A A G A T G G A T G C A A C A T Majority									
6710		6720		6730		6740		6750	
698	T	A	T	T	A	A	T	T	G
667	T	A	T	T	A	A	T	T	G
699	T	A	T	T	A	A	T	T	G
699	T	A	T	T	A	A	T	T	G
701	T	A	T	T	A	A	T	T	G
700	T	A	T	T	A	A	T	T	G
C G T C A A T A A C G A T G T T G T G A A T G G C T T A C T T A C T T A T C A A A T A G G T G A C T Majority									
6760		6770		6780		6790		6800	
748	C	G	T	C	A	A	T	A	A
517	C	G	T	C	A	A	T	A	A
749	C	G	T	C	A	A	T	A	A
749	C	G	T	C	A	A	T	A	A
751	C	G	T	C	A	A	T	A	A
750	C	G	T	C	A	A	T	A	A
A A T G A T G A T T G T G A A T A A T G G T T A T C T A G A A G G G A G A A A A A T G A A A A A G A Majority									
6810		6820		6830		6840		6850	
794	A	A	T	G	A	T	G	A	T
567	A	A	T	G	A	T	G	A	T
799	A	A	T	G	A	T	G	A	T
799	A	A	T	G	A	T	G	A	T
801	A	A	T	G	A	T	G	A	T
800	A	A	T	G	A	T	G	A	T
G A C A A A A A A T A T G G A G A G G G T T A T C A G T T A C T T T A C T A A T C C T G T C C C A A Majority									
6860		6870		6880		6890		6900	
844	G	A	C	A	A	A	A	A	T
317	G	A	C	A	A	A	A	A	T
349	G	A	C	A	A	A	A	A	T
349	G	A	C	A	A	A	A	A	T
351	G	A	C	A	A	A	A	A	T
350	G	A	C	A	A	A	A	A	T
A T T C C A T T T G G T A T A T T G G T A C A A G G T G A A A C C C A A G A T A C C A A T C A A G C Majority									
6910		6920		6930		6940		6950	
894	A	T	T	C	C	A	T	T	T
667	A	T	T	C	C	A	T	T	T
899	A	T	T	C	C	A	T	T	T
899	A	T	T	C	C	A	T	T	T
901	A	T	T	C	C	A	T	T	T
900	A	T	T	C	C	A	T	T	T
A C T T G G A A A A G T A A T T G T T A A A A A A A C G G G A G A C A A T G C T A C A C C A T T A C Majority									
6960		6970		6980		6990		7000	
944	A	C	T	T	G	G	A	A	A
917	A	C	T	T	G	G	A	A	A
949	A	C	T	T	G	G	A	A	A
949	A	C	T	T	G	G	A	A	A
951	A	C	T	T	G	G	A	A	A
950	A	C	T	T	G	G	A	A	A

FIGURE 18 S

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Alignment Report of Al-1 alignment, using J. Hein method with Weighted residue weight table.  
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G C A A A G C G A C T T T T G T G T T A A A A A A T G A C A A T G A T A A G T C A G A A A C A A G T Majority									
7010		7020		7030		7040		7050	
5994	G C A A A G C G A C T T T T G T G T T A A A A A A T G A C A A T G A T A A G T C A G A A A C A A G T 2603_all.seq								
5767	G C A A A G C G A C T T T T G T G T T A A A A A A T G A C A A T G A T A A G T C A G A A A C A A G T 18rs21_all.seq								
5999	G C A A A G C G A C T T T T G T G T T A A A A A A T G A C A A T G A T A A G T C A G A A A C A A G T coh1_all.seq								
5999	G C A A A G C G A C T T T T G T G T T A A A A A A T G A C A A T G A T A A G T C A G A A A C A A G T cjb111_all.seq								
7001	G C A A A G C G A C T T T T G T G T T A A A A A A T G A C A A T G A T A A G T C A G A A A C A A G T nem316_all.seq								
7000	G C A A A G C G A C T T T T G T G T T A A A A A A T G A C A A T G A T A A G T C A G A A A C A A G T a909_all.seq								
C A C G A A A C G G T A G A G G G T T C T G G A G A A G C A A C C T T T G A A A A C A T A A A A C C Majority									
7060		7070		7080		7090		7100	
7044	C A C G A A A C G G T A G A G G G T T C T G G A G A A G C A A C C T T T G A A A A C A T A A A A C C 2603_all.seq								
5817	C A C G A A A C G G T A G A G G G T T C T G G A G A A G C A A C C T T T G A A A A C A T A A A A C C 18rs21_all.seq								
7049	C A C G A A A C G G T A G A G G G T T C T G G A G A A G C A A C C T T T G A A A A C A T A A A A C C coh1_all.seq								
7049	C A C G A A A C G G T A G A G G G T T C T G G A G A A G C A A C C T T T G A A A A C A T A A A A C C cjb111_all.seq								
7051	C A C G A A A C G G T A G A G G G T T C T G G A G A A G C A A C C T T T G A A A A C A T A A A A C C nem316_all.seq								
7050	C A C G A A A C G G T A G A G G G T T C T G G A G A A G C A A C C T T T G A A A A C A T A A A A C C a909_all.seq								
T G G A G A C T A C A C A T T A A G A G A A G A A A C A G C A C C A A T T G G T T A T A A A A A A A Majority									
7110		7120		7130		7140		7150	
7094	T G G A G A C T A C A C A T T A A G A G A A G A A A C A G C A C C A A T T G G T T A T A A A A A A A 2603_all.seq								
5867	T G G A G A C T A C A C A T T A A G A G A A G A A A C A G C A C C A A T T G G T T A T A A A A A A A 18rs21_all.seq								
7099	T G G A G A C T A C A C A T T A A G A G A A G A A A C A G C A C C A A T T G G T T A T A A A A A A A coh1_all.seq								
7099	T G G A G A C T A C A C A T T A A G A G A A G A A A C A G C A C C A A T T G G T T A T A A A A A A A cjb111_all.seq								
7101	T G G A G A C T A C A C A T T A A G A G A A G A A A C A G C A C C A A T T G G T T A T A A A A A A A nem316_all.seq								
7100	T G G A G A C T A C A C A T T A A G A G A A G A A A C A G C A C C A A T T G G T T A T A A A A A A A a909_all.seq								
C T G A T A A A A C C T G G A A A G T T A A A G T T G C A G A T A A C G G A G C A A C A A T A A T C Majority									
7160		7170		7180		7190		7200	
7144	C T G A T A A A A C C T G G A A A G T T A A A G T T G C A G A T A A C G G A G C A A C A A T A A T C 2603_all.seq								
5917	C T G A T A A A A C C T G G A A A G T T A A A G T T G C A G A T A A C G G A G C A A C A A T A A T C 18rs21_all.seq								
7149	C T G A T A A A A C C T G G A A A G T T A A A G T T G C A G A T A A C G G A G C A A C A A T A A T C coh1_all.seq								
7149	C T G A T A A A A C C T G G A A A G T T A A A G T T G C A G A T A A C G G A G C A A C A A T A A T C cjb111_all.seq								
7151	C T G A T A A A A C C T G G A A A G T T A A A G T T G C A G A T A A C G G A G C A A C A A T A A T C nem316_all.seq								
7150	C T G A T A A A A C C T G G A A A G T T A A A G T T G C A G A T A A C G G A G C A A C A A T A A T C a909_all.seq								
G A G G G T A T G G A T G C A G A T A A A G C A G A G A A A C G A A A A G A A G T T T T G A A T G C Majority									
7210		7220		7230		7240		7250	
7194	G A G G G T A T G G A T G C A G A T A A A G C A G A G A A A C G A A A A G A A G T T T T G A A T G C 2603_all.seq								
5967	G A G G G T A T G G A T G C A G A T A A A G C A G A G A A A C G A A A A G A A G T T T T G A A T G C 18rs21_all.seq								
7199	G A G G G T A T G G A T G C A G A T A A A G C A G A G A A A C G A A A A G A A G T T T T G A A T G C coh1_all.seq								
7199	G A G G G T A T G G A T G C A G A T A A A G C A G A G A A A C G A A A A G A A G T T T T G A A T G C cjb111_all.seq								
7201	G A G G G T A T G G A T G C A G A T A A A G C A G A G A A A C G A A A A G A A G T T T T G A A T G C nem316_all.seq								
7200	G A G G G T A T G G A T G C A G A T A A A G C A G A G A A A C G A A A A G A A G T T T T G A A T G C a909_all.seq								
C C A A T A T C C A A A A T C A G C T A T T T A T G A G G A T A C A A A A G A A A A T T A C C C A T Majority									
7260		7270		7280		7290		7300	
7244	C C A A T A T C C A A A A T C A G C T A T T T A T G A G G A T A C A A A A G A A A A T T A C C C A T 2603_all.seq								
7017	C C A A T A T C C A A A A T C A G C T A T T T A T G A G G A T A C A A A A G A A A A T T A C C C A T 18rs21_all.seq								
7249	C C A A T A T C C A A A A T C A G C T A T T T A T G A G G A T A C A A A A G A A A A T T A C C C A T coh1_all.seq								
7249	C C A A T A T C C A A A A T C A G C T A T T T A T G A G G A T A C A A A A G A A A A T T A C C C A T cjb111_all.seq								
7251	C C A A T A T C C A A A A T C A G C T A T T T A T G A G G A T A C A A A A G A A A A T T A C C C A T nem316_all.seq								
7250	C C A A T A T C C A A A A T C A G C T A T T T A T G A G G A T A C A A A A G A A A A T T A C C C A T a909_all.seq								
T A G T T A A T G T A G A G G G T T C C A A A G T T G G T G A A C A A T A C A A A G C A T T G A A T Majority									
7310		7320		7330		7340		7350	
7294	T A G T T A A T G T A G A G G G T T C C A A A G T T G G T G A A C A A T A C A A A G C A T T G A A T 2603_all.seq								
7067	T A G T T A A T G T A G A G G G T T C C A A A G T T G G T G A A C A A T A C A A A G C A T T G A A T 18rs21_all.seq								
7299	T A G T T A A T G T A G A G G G T T C C A A A G T T G G T G A A C A A T A C A A A G C A T T G A A T coh1_all.seq								
7299	T A G T T A A T G T A G A G G G T T C C A A A G T T G G T G A A C A A T A C A A A G C A T T G A A T cjb111_all.seq								
7301	T A G T T A A T G T A G A G G G T T C C A A A G T T G G T G A A C A A T A C A A A G C A T T G A A T nem316_all.seq								
7300	T A G T T A A T G T A G A G G G T T C C A A A G T T G G T G A A C A A T A C A A A G C A T T G A A T a909_all.seq								

FIGURE 18 T



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C C A A T A A A T G C G A A A A G A T G C T C G A A G A G A G A T T G C T G A A G G T T G G T T A T C Majority										
7360		7370		7380		7390		7400		
7344	C C A A T A A A T G C G A A A A G A T G C T C G A A G A G A G A T T G C T G A A G G T T G G T T A T C									2603_ail.seq
7117	C C A A T A A A T G C G A A A A G A T G C T C G A A G A G A G A T T G C T G A A G G T T G G T T A T C									18rs21_ail.seq
7349	C C A A T A A A T G C G A A A A G A T G C T C G A A G A G A G A T T G C T G A A G G T T G G T T A T C									cohl_ail.seq
7349	C C A A T A A A T G C G A A A A G A T G C T C G A A G A G A G A T T G C T G A A G G T T G G T T A T C									cjb111_ail.seq
7351	C C A A T A A A T G C G A A A A G A T G C T C G A A G A G A G A T T G C T G A A G G T T G G T T A T C									nen316_ail.seq
7350	C C A A T A A A T G C G A A A A G A T G C T C G A A G A G A G A T T G C T G A A G G T T G G T T A T C									a909_ail.seq
A A A A A A A A T T A C A G G G G T C A A T G A T C T C G A T A A G A A T A A A T A T A A A A T T G Majority										
7410		7420		7430		7440		7450		
7394	A A A A A A A A T T A C A G G G G T C A A T G A T C T C G A T A A G A A T A A A T A T A A A A T T G									2603_ail.seq
7167	A A A A A A A A T T A C A G G G G T C A A T G A T C T C G A T A A G A A T A A A T A T A A A A T T G									18rs21_ail.seq
7399	A A A A A A A A T T A C A G G G G T C A A T G A T C T C G A T A A G A A T A A A T A T A A A A T T G									cohl_ail.seq
7399	A A A A A A A A T T A C A G G G G T C A A T G A T C T C G A T A A G A A T A A A T A T A A A A T T G									cjb111_ail.seq
7401	A A A A A A A A T T A C A G G G G T C A A T G A T C T C G A T A A G A A T A A A T A T A A A A T T G									nen316_ail.seq
7400	A A A A A A A A T T A C A G G G G T C A A T G A T C T C G A T A A G A A T A A A T A T A A A A T T G									a909_ail.seq
A A T T A A C T G T T G A G G G T A A A A C C A C T G T T G A A A C G A A A G A A C T T A A T C A A Majority										
7460		7470		7480		7490		7500		
7444	A A T T A A C T G T T G A G G G T A A A A C C A C T G T T G A A A C G A A A G A A C T T A A T C A A									2603_ail.seq
7217	A A T T A A C T G T T G A G G G T A A A A C C A C T G T T G A A A C G A A A G A A C T T A A T C A A									18rs21_ail.seq
7449	A A T T A A C T G T T G A G G G T A A A A C C A C T G T T G A A A C G A A A G A A C T T A A T C A A									cohl_ail.seq
7449	A A T T A A C T G T T G A G G G T A A A A C C A C T G T T G A A A C G A A A G A A C T T A A T C A A									cjb111_ail.seq
7451	A A T T A A C T G T T G A G G G T A A A A C C A C T G T T G A A A C G A A A G A A C T T A A T C A A									nen316_ail.seq
7450	A A T T A A C T G T T G A G G G T A A A A C C A C T G T T G A A A C G A A A G A A C T T A A T C A A									a909_ail.seq
C C A C T A G A T G T C G T T G T G C T A T T A G A T A A T T C A A A T A G T A T G A A T A A T G A Majority										
7510		7520		7530		7540		7550		
7494	C C A C T A G A T G T C G T T G T G C T A T T A G A T A A T T C A A A T A G T A T G A A T A A T G A									2603_ail.seq
7267	C C A C T A G A T G T C G T T G T G C T A T T A G A T A A T T C A A A T A G T A T G A A T A A T G A									18rs21_ail.seq
7499	C C A C T A G A T G T C G T T G T G C T A T T A G A T A A T T C A A A T A G T A T G A A T A A T G A									cohl_ail.seq
7499	C C A C T A G A T G T C G T T G T G C T A T T A G A T A A T T C A A A T A G T A T G A A T A A T G A									cjb111_ail.seq
7501	C C A C T A G A T G T C G T T G T G C T A T T A G A T A A T T C A A A T A G T A T G A A T A A T G A									nen316_ail.seq
7500	C C A C T A G A T G T C G T T G T G C T A T T A G A T A A T T C A A A T A G T A T G A A T A A T G A									a909_ail.seq
A A G A G C C A A T A A T T C T C A A A G A G C A T T A A A A G C T G G G G A A G C A G T T G A A A Majority										
7560		7570		7580		7590		7600		
7544	A A G A G C C A A T A A T T C T C A A A G A G C A T T A A A A G C T G G G G A A G C A G T T G A A A									2603_ail.seq
7317	A A G A G C C A A T A A T T C T C A A A G A G C A T T A A A A G C T G G G G A A G C A G T T G A A A									18rs21_ail.seq
7549	A A G A G C C A A T A A T T C T C A A A G A G C A T T A A A A G C T G G G G A A G C A G T T G A A A									cohl_ail.seq
7549	A A G A G C C A A T A A T T C T C A A A G A G C A T T A A A A G C T G G G G A A G C A G T T G A A A									cjb111_ail.seq
7551	A A G A G C C A A T A A T T C T C A A A G A G C A T T A A A A G C T G G G G A A G C A G T T G A A A									nen316_ail.seq
7550	A A G A G C C A A T A A T T C T C A A A G A G C A T T A A A A G C T G G G G A A G C A G T T G A A A									a909_ail.seq
A G C T G A T T G A T A A A A T T A C A T C A A A T A A A G A C A A T A G A G T A G C T C T T G T G Majority										
7610		7620		7630		7640		7650		
7594	A G C T G A T T G A T A A A A T T A C A T C A A A T A A A G A C A A T A G A G T A G C T C T T G T G									2603_ail.seq
7367	A G C T G A T T G A T A A A A T T A C A T C A A A T A A A G A C A A T A G A G T A G C T C T T G T G									18rs21_ail.seq
7599	A G C T G A T T G A T A A A A T T A C A T C A A A T A A A G A C A A T A G A G T A G C T C T T G T G									cohl_ail.seq
7599	A G C T G A T T G A T A A A A T T A C A T C A A A T A A A G A C A A T A G A G T A G C T C T T G T G									cjb111_ail.seq
7601	A G C T G A T T G A T A A A A T T A C A T C A A A T A A A G A C A A T A G A G T A G C T C T T G T G									nen316_ail.seq
7600	A G C T G A T T G A T A A A A T T A C A T C A A A T A A A G A C A A T A G A G T A G C T C T T G T G									a909_ail.seq
A C A T A T G C C T C A A C C A T T T T T G A T G G T A C T G A A G C G A C C G T A T C A A A G G G Majority										
7660		7670		7680		7690		7700		
7644	A C A T A T G C C T C A A C C A T T T T T G A T G G T A C T G A A G C G A C C G T A T C A A A G G G									2603_ail.seq
7417	A C A T A T G C C T C A A C C A T T T T T G A T G G T A C T G A A G C G A C C G T A T C A A A G G G									18rs21_ail.seq
7649	A C A T A T G C C T C A A C C A T T T T T G A T G G T A C T G A A G C G A C C G T A T C A A A G G G									cohl_ail.seq
7649	A C A T A T G C C T C A A C C A T T T T T G A T G G T A C T G A A G C G A C C G T A T C A A A G G G									cjb111_ail.seq
7651	A C A T A T G C C T C A A C C A T T T T T G A T G G T A C T G A A G C G A C C G T A T C A A A G G G									nen316_ail.seq
7650	A C A T A T G C C T C A A C C A T T T T T G A T G G T A C T G A A G C G A C C G T A T C A A A G G G									a909_ail.seq

FIGURE 18 U

A G T T G C C G A T C A A A A T G G T A A A G C G C T G A A T G A T A G T G T A T C A T G G G A T T Majority									
7710		7720		7730		7740		7750	
7694	A	G	T	T	G	C	C	G	A
7697	A	G	T	T	G	C	C	G	A
7699	A	G	T	T	G	C	C	G	A
7699	A	G	T	T	G	C	C	G	A
7701	A	G	T	T	G	C	C	G	A
7700	A	G	T	T	G	C	C	G	A
A T C A T A A A A C T A C T T T T A C A G C A A C T A C A C A T A A T T A C A G T T A T T T A A A T Majority									
7760		7770		7780		7790		7800	
7744	A	T	C	A	T	A	A	A	C
7751	A	T	C	A	T	A	A	A	C
7749	A	T	C	A	T	A	A	A	C
7749	A	T	C	A	T	A	A	A	C
7751	A	T	C	A	T	A	A	A	C
7750	A	T	C	A	T	A	A	A	C
T T A A C A A A T G A T G C T A A C G A A G T T A A T A T T C T A A A G T C A A G A A T T C C A A A Majority									
7810		7820		7830		7840		7850	
7794	T	T	A	A	C	A	A	A	T
7797	T	T	A	A	C	A	A	A	T
7799	T	T	A	A	C	A	A	A	T
7799	T	T	A	A	C	A	A	A	T
7801	T	T	A	A	C	A	A	A	T
7800	T	T	A	A	C	A	A	A	T
G G A A G C C G A G C A T A T A A A T G G G G A T C G C A C G C T C T A T C A A T T T G G T G C G A Majority									
7860		7870		7880		7890		7900	
7844	G	G	A	A	G	C	C	G	A
7847	G	G	A	A	G	C	C	G	A
7849	G	G	A	A	G	C	C	G	A
7849	G	G	A	A	G	C	C	G	A
7851	G	G	A	A	G	C	C	G	A
7850	G	G	A	A	G	C	C	G	A
C A T T T A C T C A A A A A G C T C T A A T G A A A G C A A A T G A A A T T T T A G A G A C A C A A Majority									
7910		7920		7930		7940		7950	
7894	C	A	T	T	A	C	T	C	A
7897	C	A	T	T	A	C	T	C	A
7899	C	A	T	T	A	C	T	C	A
7899	C	A	T	T	A	C	T	C	A
7901	C	A	T	T	A	C	T	C	A
7900	C	A	T	T	A	C	T	C	A
A G T T C T A A T G C T A G A A A A A A C T T A T T T T T C A C G T A A C T G A T G G T G T C C C Majority									
7960		7970		7980		7990		8000	
7944	A	G	T	T	C	T	A	A	T
7947	A	G	T	T	C	T	A	A	T
7949	A	G	T	T	C	T	A	A	T
7949	A	G	T	T	C	T	A	A	T
7951	A	G	T	T	C	T	A	A	T
7950	A	G	T	T	C	T	A	A	T
T A C G A T G T C T T A T G C C A T A A A T T T T A A T C C T T A T A T A T C A A C A T C T T A C C Majority									
8010		8020		8030		8040		8050	
7994	T	A	C	G	A	T	G	T	C
7997	T	A	C	G	A	T	G	T	C
7999	T	A	C	G	A	T	G	T	C
7999	T	A	C	G	A	T	G	T	C
8001	T	A	C	G	A	T	G	T	C
8000	T	A	C	G	A	T	G	T	C

FIGURE 18 V



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Alignment Report of Al-1 alignment, using J. Hein method with Weighted residue weight table.  
Thursday, July 29, 2004 5:46 PM

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A A A A C C A G G T T T A A T T C T T T T T T A A A T A A A A T A C C A G A T A G A A G T G G T A T T Majority									
8060		8070		8080		8090		8100	
1044	A A A A C C A G T T T A A T T C T T T T T T A A A T A A A A T A C C A G A T A G A A G T G G T A T T 2603_all.seq								
1817	A A A A C C A G T T T A A T T C T T T T T T A A A T A A A A T A C C A G A T A G A A G T G G T A T T 18rs21_all.seq								
1049	A A A A C C A G T T T A A T T C T T T T T T A A A T A A A A T A C C A G A T A G A A G T G G T A T T coh1_all.seq								
1049	A A A A C C A G T T T A A T T C T T T T T T A A A T A A A A T A C C A G A T A G A A G T G G T A T T cjb111_all.seq								
1051	A A A A C C A G T T T A A T T C T T T T T T A A A T A A A A T A C C A G A T A G A A G T G G T A T T nem316_all.seq								
1050	A A A A C C A G T T T A A T T C T T T T T T A A A T A A A A T A C C A G A T A G A A G T G G T A T T a909_all.seq								
C T C C A A G A G G A T T T T A T A A T C A A T G G T G A T G A T T A T C A A A T A G T A A A A G G Majority									
8110		8120		8130		8140		8150	
1094	C T C C A A G A G G A T T T T A T A A T C A A T G G T G A T G A T T A T C A A A T A G T A A A A G G 2603_all.seq								
1867	C T C C A A G A G G A T T T T A T A A T C A A T G G T G A T G A T T A T C A A A T A G T A A A A G G 18rs21_all.seq								
1099	C T C C A A G A G G A T T T T A T A A T C A A T G G T G A T G A T T A T C A A A T A G T A A A A G G coh1_all.seq								
1099	C T C C A A G A G G A T T T T A T A A T C A A T G G T G A T G A T T A T C A A A T A G T A A A A G G cjb111_all.seq								
1101	C T C C A A G A G G A T T T T A T A A T C A A T G G T G A T G A T T A T C A A A T A G T A A A A G G nem316_all.seq								
1100	C T C C A A G A G G A T T T T A T A A T C A A T G G T G A T G A T T A T C A A A T A G T A A A A G G a909_all.seq								
A G A T G G A G A G A G T T T T A A A C T G T T T T C G G A T A G A A A A G T T C C T G T T A C T G Majority									
8160		8170		8180		8190		8200	
1144	A G A T G G A G A G A G T T T T A A A C T G T T T T C G G A T A G A A A A G T T C C T G T T A C T G 2603_all.seq								
1917	A G A T G G A G A G A G T T T T A A A C T G T T T T C G G A T A G A A A A G T T C C T G T T A C T G 18rs21_all.seq								
1149	A G A T G G A G A G A G T T T T A A A C T G T T T T C G G A T A G A A A A G T T C C T G T T A C T G coh1_all.seq								
1149	A G A T G G A G A G A G T T T T A A A C T G T T T T C G G A T A G A A A A G T T C C T G T T A C T G cjb111_all.seq								
1151	A G A T G G A G A G A G T T T T A A A C T G T T T T C G G A T A G A A A A G T T C C T G T T A C T G nem316_all.seq								
1150	A G A T G G A G A G A G T T T T A A A C T G T T T T C G G A T A G A A A A G T T C C T G T T A C T G a909_all.seq								
G A G G A A C G A C A C A A G C A G C T T A T C G A G T A C C G C A A A A T C A A C T C T C T G T A Majority									
8210		8220		8230		8240		8250	
194	G A G G A A C G A C A C A A G C A G C T T A T C G A G T A C C G C A A A A T C A A C T C T C T G T A 2603_all.seq								
967	G A G G A A C G A C A C A A G C A G C T T A T C G A G T A C C G C A A A A T C A A C T C T C T G T A 18rs21_all.seq								
199	G A G G A A C G A C A C A A G C A G C T T A T C G A G T A C C G C A A A A T C A A C T C T C T G T A coh1_all.seq								
199	G A G G A A C G A C A C A A G C A G C T T A T C G A G T A C C G C A A A A T C A A C T C T C T G T A cjb111_all.seq								
201	G A G G A A C G A C A C A A G C A G C T T A T C G A G T A C C G C A A A A T C A A C T C T C T G T A nem316_all.seq								
200	G A G G A A C G A C A C A A G C A G C T T A T C G A G T A C C G C A A A A T C A A C T C T C T G T A a909_all.seq								
A T G A G T A A T G A G G G A T A T G C A A T T A A T A G T G G A T A T A T T T A T C T C T A T T G Majority									
8260		8270		8280		8290		8300	
244	A T G A G T A A T G A G G G A T A T G C A A T T A A T A G T G G A T A T A T T T A T C T C T A T T G 2603_all.seq								
017	A T G A G T A A T G A G G G A T A T G C A A T T A A T A G T G G A T A T A T T T A T C T C T A T T G 18rs21_all.seq								
249	A T G A G T A A T G A G G G A T A T G C A A T T A A T A G T G G A T A T A T T T A T C T C T A T T G coh1_all.seq								
249	A T G A G T A A T G A G G G A T A T G C A A T T A A T A G T G G A T A T A T T T A T C T C T A T T G cjb111_all.seq								
251	A T G A G T A A T G A G G G A T A T G C A A T T A A T A G T G G A T A T A T T T A T C T C T A T T G nem316_all.seq								
250	A T G A G T A A T G A G G G A T A T G C A A T T A A T A G T G G A T A T A T T T A T C T C T A T T G a909_all.seq								
G A G A G A T T A C A A C T G G G T C T A T C C A T T T G A T C C T A A G A C A A A G A A A G T T T Majority									
8310		8320		8330		8340		8350	
294	G A G A G A T T A C A A C T G G G T C T A T C C A T T T G A T C C T A A G A C A A A G A A A G T T T 2603_all.seq								
067	G A G A G A T T A C A A C T G G G T C T A T C C A T T T G A T C C T A A G A C A A A G A A A G T T T 18rs21_all.seq								
299	G A G A G A T T A C A A C T G G G T C T A T C C A T T T G A T C C T A A G A C A A A G A A A G T T T coh1_all.seq								
299	G A G A G A T T A C A A C T G G G T C T A T C C A T T T G A T C C T A A G A C A A A G A A A G T T T cjb111_all.seq								
301	G A G A G A T T A C A A C T G G G T C T A T C C A T T T G A T C C T A A G A C A A A G A A A G T T T nem316_all.seq								
300	G A G A G A T T A C A A C T G G G T C T A T C C A T T T G A T C C T A A G A C A A A G A A A G T T T a909_all.seq								
C T G C A A C G A A A G A A A T C A A A A C T C A T G G T G A G C C A A C A A C A T T A T A C T T T Majority									
8360		8370		8380		8390		8400	
344	C T G C A A C G A A A G A A A T C A A A A C T C A T G G T G A G C C A A C A A C A T T A T A C T T T 2603_all.seq								
117	C T G C A A C G A A A G A A A T C A A A A C T C A T G G T G A G C C A A C A A C A T T A T A C T T T 18rs21_all.seq								
349	C T G C A A C G A A A G A A A T C A A A A C T C A T G G T G A G C C A A C A A C A T T A T A C T T T coh1_all.seq								
349	C T G C A A C G A A A G A A A T C A A A A C T C A T G G T G A G C C A A C A A C A T T A T A C T T T cjb111_all.seq								
351	C T G C A A C G A A A G A A A T C A A A A C T C A T G G T G A G C C A A C A A C A T T A T A C T T T nem316_all.seq								
350	C T G C A A C G A A A G A A A T C A A A A C T C A T G G T G A G C C A A C A A C A T T A T A C T T T a909_all.seq								

FIGURE 18 W

**FIGURE 18 X**



A A A C A G T G A T G C G C G A A T T T T A A A A G A T G T T A C A G T G A C T T A T G A T A A G A Majority										
8760		8770		8780		8790		8800		
8744	A A A C A G T G A T G C G C G A A T T T T A A A A G A T G T T A C A G T G A C T T A T G A T A A G A									2603_all.seq
8517	A A A C A G T G A T G C G C G A A T T T T A A A A G A T G T T A C A G T G A C T T A T G A T A A G A									18rs21_all.seq
8749	A A A C A G T G A T G C G C G A A T T T T A A A A G A T G T T A C A G T G A C T T A T G A T A A G A									cohl_all.seq
8749	A A A C A G T G A T G C G C G A A T T T T A A A A G A T G T T A C A G T G A C T T A T G A T A A G A									cjb111_all.seq
8751	A A A C A G T G A T G C G C G A A T T T T A A A A G A T G T T A C A G T G A C T T A T G A T A A G A									nem316_all.seq
8750	A A A C A G T G A T G C G C G A A T T T T A A A A G A T G T T A C A G T G A C T T A T G A T A A G A									a909_all.seq
C A T C T C A A A C C A T C A A A A T C A A T C A T T T G A A C T T A G G A A G T G G A C A A A A A Majority										
8810		8820		8830		8840		8850		
8794	C A T C T C A A A C C A T C A A A A T C A A T C A T T T G A A C T T A G G A A G T G G A C A A A A A									2603_all.seq
8567	C A T C T C A A A C C A T C A A A A T C A A T C A T T T G A A C T T A G G A A G T G G A C A A A A A									18rs21_all.seq
8799	C A T C T C A A A C C A T C A A A A T C A A T C A T T T G A A C T T A G G A A G T G G A C A A A A A									cohl_all.seq
8799	C A T C T C A A A C C A T C A A A A T C A A T C A T T T G A A C T T A G G A A G T G G A C A A A A A									cjb111_all.seq
8801	C A T C T C A A A C C A T C A A A A T C A A T C A T T T G A A C T T A G G A A G T G G A C A A A A A									nem316_all.seq
8800	C A T C T C A A A C C A T C A A A A T C A A T C A T T T G A A C T T A G G A A G T G G A C A A A A A									a909_all.seq
G T A G T T C T T A C C T A T G A T G T A C G T T T A A A A G A T A A C T A T A T A A G T A A C A A Majority										
8860		8870		8880		8890		8900		
8844	G T A G T T C T T A C C T A T G A T G T A C G T T T A A A A G A T A A C T A T A T A A A G T A A C A A									2603_all.seq
8617	G T A G T T C T T A C C T A T G A T G T A C G T T T A A A A G A T A A C T A T A T A A A G T A A C A A									18rs21_all.seq
8849	G T A G T T C T T A C C T A T G A T G T A C G T T T A A A A G A T A A C T A T A T A A A G T A A C A A									cohl_all.seq
8849	G T A G T T C T T A C C T A T G A T G T A C G T T T A A A A G A T A A C T A T A T A A A G T A A C A A									cjb111_all.seq
8851	G T A G T T C T T A C C T A T G A T G T A C G T T T A A A A G A T A A C T A T A T A A A G T A A C A A									nem316_all.seq
8850	G T A G T T C T T A C C T A T G A T G T A C G T T T A A A A G A T A A C T A T A T A A A G T A A C A A									a909_all.seq
A T T T T A C A A T A C A A A T A A T C G T A C A A C G C T A A G T C C G A A G A G T G A A A A A G Majority										
8910		8920		8930		8940		8950		
8894	A T T T T A C A A T A C A A A T A A T C G T A C A A C G C T A A G T C C G A A G A G T G A A A A A G									2603_all.seq
8667	A T T T T A C A A T A C A A A T A A T C G T A C A A C G C T A A G T C C G A A G A G T G A A A A A G									18rs21_all.seq
8899	A T T T T A C A A T A C A A A T A A T C G T A C A A C G C T A A G T C C G A A G A G T G A A A A A G									cohl_all.seq
8899	A T T T T A C A A T A C A A A T A A T C G T A C A A C G C T A A G T C C G A A G A G T G A A A A A G									cjb111_all.seq
8901	A T T T T A C A A T A C A A A T A A T C G T A C A A C G C T A A G T C C G A A G A G T G A A A A A G									nem316_all.seq
8900	A T T T T A C A A T A C A A A T A A T C G T A C A A C G C T A A G T C C G A A G A G T G A A A A A G									a909_all.seq
A A C C A A A T A C T A T T C G T G A T T T C C C A A T T C C C A A A A T T C G T G A T G T T C G T Majority										
8960		8970		8980		8990		9000		
8944	A A C C A A A T A C T A T T C G T G A T T T C C C A A T T C C C A A A A T T C G T G A T G T T C G T									2603_all.seq
8717	A A C C A A A T A C T A T T C G T G A T T T C C C A A T T C C C A A A A T T C G T G A T G T T C G T									18rs21_all.seq
8949	A A C C A A A T A C T A T T C G T G A T T T C C C A A T T C C C A A A A T T C G T G A T G T T C G T									cohl_all.seq
8949	A A C C A A A T A C T A T T C G T G A T T T C C C A A T T C C C A A A A T T C G T G A T G T T C G T									cjb111_all.seq
8951	A A C C A A A T A C T A T T C G T G A T T T C C C A A T T C C C A A A A T T C G T G A T G T T C G T									nem316_all.seq
8950	A A C C A A A T A C T A T T C G T G A T T T C C C A A T T C C C A A A A T T C G T G A T G T T C G T									a909_all.seq
G A G T T T C C G G T A C T A A C C A T C A G T A A T C A G A A G A A A A T G G G T G A G G T T G A Majority										
9010		9020		9030		9040		9050		
8994	G A G T T T C C G G T A C T A A C C A T C A G T A A T C A G A A G A A A A T G G G T G A G G T T G A									2603_all.seq
8767	G A G T T T C C G G T A C T A A C C A T C A G T A A T C A G A A G A A A A T G G G T G A G G T T G A									18rs21_all.seq
8999	G A G T T T C C G G T A C T A A C C A T C A G T A A T C A G A A G A A A A T G G G T G A G G T T G A									cohl_all.seq
8999	G A G T T T C C G G T A C T A A C C A T C A G T A A T C A G A A G A A A A T G G G T G A G G T T G A									cjb111_all.seq
8001	G A G T T T C C G G T A C T A A C C A T C A G T A A T C A G A A G A A A A T G G G T G A G G T T G A									nem316_all.seq
8000	G A G T T T C C G G T A C T A A C C A T C A G T A A T C A G A A G A A A A T G G G T G A G G T T G A									a909_all.seq
A T T T A T T A A A G T T A A T A A A G A C A A A C A T T C A G A A T C G C T T T T G G G A G C T A Majority										
9060		9070		9080		9090		9100		
8044	A T T T A T T A A A G T T A A T A A A G A C A A A C A T T C A G A A T C G C T T T T G G G A G C T A									2603_all.seq
8817	A T T T A T T A A A G T T A A T A A A G A C A A A C A T T C A G A A T C G C T T T T G G G A G C T A									18rs21_all.seq
8049	A T T T A T T A A A G T T A A T A A A G A C A A A C A T T C A G A A T C G C T T T T G G G A G C T A									cohl_all.seq
8049	A T T T A T T A A A G T T A A T A A A G A C A A A C A T T C A G A A T C G C T T T T G G G A G C T A									cjb111_all.seq
8051	A T T T A T T A A A G T T A A T A A A G A C A A A C A T T C A G A A T C G C T T T T G G G A G C T A									nem316_all.seq
8050	A T T T A T T A A A G T T A A T A A A G A C A A A C A T T C A G A A T C G C T T T T G G G A G C T A									a909_all.seq

FIGURE 18 Y

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A G T T T C A A C T T C A G A T A G A A A A A G A T T T T C T G G G T A T A A G C A A T T T G T T Majority																																																	
9110										9120										9130										9140										9150									
9094	A G T T T C A A C T T C A G A T A G A A A A A G A T T T T T C T G G G T A T A A G C A A T T T G T T																														2603_all.seq																		
8867	A G T T T C A A C T T C A G A T A G A A A A A G A T T T T T C T G G G T A T A A G C A A T T T G T T																														18rs21_all.seq																		
9099	A G T T T C A A C T T C A G A T A G A A A A A G A T T T T T C T G G G T A T A A G C A A T T T G T T																														cohl_all.seq																		
9099	A G T T T C A A C T T C A G A T A G A A A A A G A T T T T T C T G G G T A T A A G C A A T T T G T T																														cjb111_all.seq																		
9101	A G T T T C A A C T T C A G A T A G A A A A A G A T T T T T C T G G G T A T A A G C A A T T T G T T																														nen316_all.seq																		
9100	A G T T T C A A C T T C A G A T A G A A A A A G A T T T T T C T G G G T A T A A G C A A T T T G T T																														a909_all.seq																		
C C A G A G G G A A G T G A T G T T A C A A C A A A G A A T G A T G G T A A A A T T T A T T T T A A Majority																																																	
9160										9170										9180										9190										9200									
9144	C C A G A G G G A A G T G A T G T T A C A A C A A A G A A T G A T G G T A A A A T T T A T T T T A A																														2603_all.seq																		
8917	C C A G A G G G A A G T G A T G T T A C A A C A A A G A A T G A T G G T A A A A T T T A T T T T A A																														18rs21_all.seq																		
9149	C C A G A G G G A A G T G A T G T T A C A A C A A A G A A T G A T G G T A A A A T T T A T T T T A A																														cohl_all.seq																		
9149	C C A G A G G G A A G T G A T G T T A C A A C A A A G A A T G A T G G T A A A A T T T A T T T T A A																														cjb111_all.seq																		
9151	C C A G A G G G A A G T G A T G T T A C A A C A A A G A A T G A T G G T A A A A T T T A T T T T A A																														nen316_all.seq																		
9150	C C A G A G G G A A G T G A T G T T A C A A C A A A G A A T G A T G G T A A A A T T T A T T T T A A																														a909_all.seq																		
A G C A C T T C A A G A T G G T A A C T A T A A A T T A T A T G A A A T T T C A A G T C C A G A T G Majority																																																	
9210										9220										9230										9240										9250									
9194	A G C A C T T C A A G A T G G T A A C T A T A A A T T A T A T G A A A T T T C A A G T C C A G A T G																														2603_all.seq																		
9197	A G C A C T T C A A G A T G G T A A C T A T A A A T T A T A T G A A A T T T C A A G T C C A G A T G																														18rs21_all.seq																		
9199	A G C A C T T C A A G A T G G T A A C T A T A A A T T A T A T G A A A T T T C A A G T C C A G A T G																														cohl_all.seq																		
9199	A G C A C T T C A A G A T G G T A A C T A T A A A T T A T A T G A A A T T T C A A G T C C A G A T G																														cjb111_all.seq																		
9201	A G C A C T T C A A G A T G G T A A C T A T A A A T T A T A T G A A A T T T C A A G T C C A G A T G																														nen316_all.seq																		
9200	A G C A C T T C A A G A T G G T A A C T A T A A A T T A T A T G A A A T T T C A A G T C C A G A T G																														a909_all.seq																		
G C T A T A T A G A G G T T A A A A C G A A A C C T G T T G T G A C A T T T A C A A T T C A A A A T Majority																																																	
9260										9270										9280										9290										9300									
9244	G C T A T A T A G A G G T T A A A A C G A A A C C T G T T G T G A C A T T T A C A A T T C A A A A T																														2603_all.seq																		
9247	G C T A T A T A G A G G T T A A A A C G A A A C C T G T T G T G A C A T T T A C A A T T C A A A A T																														18rs21_all.seq																		
9249	G C T A T A T A G A G G T T A A A A C G A A A C C T G T T G T G A C A T T T A C A A T T C A A A A T																														cohl_all.seq																		
9249	G C T A T A T A G A G G T T A A A A C G A A A C C T G T T G T G A C A T T T A C A A T T C A A A A T																														cjb111_all.seq																		
9251	G C T A T A T A G A G G T T A A A A C G A A A C C T G T T G T G A C A T T T A C A A T T C A A A A T																														nen316_all.seq																		
9250	G C T A T A T A G A G G T T A A A A C G A A A C C T G T T G T G A C A T T T A C A A T T C A A A A T																														a909_all.seq																		
G G A G A A G T T A C G A A A C C T G A A A G C A G A T C C A A A T G C T A A T A A A A A T C A A A T Majority																																																	
9310										9320										9330										9340										9350									
9294	G G A G A A G T T A C G A A A C C T G A A A G C A G A T C C A A A T G C T A A T A A A A A T C A A A T																														2603_all.seq																		
9297	G G A G A A G T T A C G A A A C C T G A A A G C A G A T C C A A A T G C T A A T A A A A A T C A A A T																														18rs21_all.seq																		
9299	G G A G A A G T T A C G A A A C C T G A A A G C A G A T C C A A A T G C T A A T A A A A A T C A A A T																														cohl_all.seq																		
9299	G G A G A A G T T A C G A A A C C T G A A A G C A G A T C C A A A T G C T A A T A A A A A T C A A A T																														cjb111_all.seq																		
9301	G G A G A A G T T A C G A A A C C T G A A A G C A G A T C C A A A T G C T A A T A A A A A T C A A A T																														nen316_all.seq																		
9300	G G A G A A G T T A C G A A A C C T G A A A G C A G A T C C A A A T G C T A A T A A A A A T C A A A T																														a909_all.seq																		
C G G G T A T C T T G A A G G A A A T G G T A A A C A T C T T A T T A C C A A C A C T C C C A A A C Majority																																																	
9360										9370										9380										9390										9400									
9344	C G G G T A T C T T G A A G G A A A T G G T A A A C A T C T T A T T A C C A A C A C T C C C A A A C																														2603_all.seq																		
9347	C G G G T A T C T T G A A G G A A A T G G T A A A C A T C T T A T T A C C A A C A C T C C C A A A C																														18rs21_all.seq																		
9349	C G G G T A T C T T G A A G G A A A T G G T A A A C A T C T T A T T A C C A A C A C T C C C A A A C																														cohl_all.seq																		
9349	C G G G T A T C T T G A A G G A A A T G G T A A A C A T C T T A T T A C C A A C A C T C C C A A A C																														cjb111_all.seq																		
9351	C G G G T A T C T T G A A G G A A A T G G T A A A C A T C T T A T T A C C A A C A C T C C C A A A C																														nen316_all.seq																		
9350	C G G G T A T C T T G A A G G A A A T G G T A A A C A T C T T A T T A C C A A C A C T C C C A A A C																														a909_all.seq																		
G C C C A C C A G G T G T T T T T C C T A A A A C A G G G G G A A T T G G T A C A A T T G T C T A T Majority																																																	
9410										9420										9430										9440										9450									
9394	G C C C A C C A G G T G T T T T T C C T A A A A C A G G G G G A A T T G G T A C A A T T G T C T A T																														2603_all.seq																		
9397	G C C C A C C A G G T G T T T T T C C T A A A A C A G G G G G A A T T G G T A C A A T T G T C T A T																														18rs21_all.seq																		
9399	G C C C A C C A G G T G T T T T T C C T A A A A C A G G G G G A A T T G G T A C A A T T G T C T A T																														cohl_all.seq																		
9399	G C C C A C C A G G T G T T T T T C C T A A A A C A G G G G G A A T T G G T A C A A T T G T C T A T																														cjb111_all.seq																		
9401	G C C C A C C A G G T G T T T T T C C T A A A A C A G G G G G A A T T G G T A C A A T T G T C T A T																														nen316_all.seq																		
9400	G C C C A C C A G G T G T T T T T C C T A A A A C A G G G G G A A T T G G T A C A A T T G T C T A T																														a909_all.seq																		



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A T A T T A G T T G G T T C T A C T T T T A T G A T A C T T A C C A T T T G T T C T T T C C G T C G Majority									
9460		9470		9480		9490		9500	
9444	A	T	A	T	T	A	G	T	T
9217	A	T	A	T	T	A	G	T	T
9449	A	T	A	T	T	A	G	T	T
9449	A	T	A	T	T	A	G	T	T
9451	A	T	A	T	T	A	G	T	T
9450	A	T	A	T	T	A	G	T	T
T A A A C A A T T G T A A G G T G T C G T T G A A A T T A T T A A A T A A T A G A A A A T G A C T A Majority									
9510		9520		9530		9540		9550	
9494	T	A	A	A	C	A	A	T	T
9267	T	A	A	A	C	A	A	T	T
9499	T	A	A	A	C	A	A	T	T
9499	T	A	A	A	C	A	A	T	T
9501	T	A	A	A	C	A	A	T	T
9500	T	A	A	A	C	A	A	T	T
G T T T T G G C C T T T C C C T A T T G T C A G T C A G A T T A G T T A T T A T C A A G C T T C T C Majority									
9560		9570		9580		9590		9600	
9544	G	T	T	T	G	G	C	C	T
9317	G	T	T	T	G	G	C	C	T
9549	G	T	T	T	G	G	C	C	T
9549	G	T	T	T	G	G	C	C	T
9551	G	T	T	T	G	G	C	C	T
9550	G	T	T	T	G	G	C	C	T
A T G C C A A T A T T A A T G C C T T T A A A A G A A C A G T T A C A A C C A T T G A C C G T A C G Majority									
9610		9620		9630		9640		9650	
594	A	T	G	C	C	A	A	T	A
367	A	T	G	C	C	A	A	T	A
599	A	T	G	C	C	A	A	T	A
599	A	T	G	C	C	A	A	T	A
601	A	T	G	C	C	A	A	T	A
600	A	T	G	C	C	A	A	T	A
G A G A T T C A A C G A C G G T T A G G T C T G G C T A A A G C C T A C A A T G C T A G T A T T T C Majority									
9660		9670		9680		9690		9700	
644	G	A	G	A	T	T	C	A	A
417	G	A	G	A	T	T	C	A	A
649	G	A	G	A	T	T	C	A	A
649	G	A	G	A	T	T	C	A	A
651	G	A	G	A	T	T	C	A	A
650	G	A	G	A	T	T	C	A	A
T G G A A C A A G T A G T C A G T C G A C T C A A T C T G T G C T G A G A G A T T C T T A T T C T G Majority									
9710		9720		9730		9740		9750	
394	T	G	G	A	A	C	A	A	G
467	T	G	G	A	A	C	A	A	G
399	T	G	G	A	A	C	A	A	G
399	T	G	G	A	A	C	A	A	G
701	T	G	G	A	A	C	A	A	G
700	T	G	G	A	A	C	A	A	G
A G G A G C A A A A A A G G C A G G G T G G A C T G A A T A C G C T A G G A T G T T A G A A G T C A Majority									
9760		9770		9780		9790		9800	
744	A	G	G	A	G	C	A	A	A
517	A	G	G	A	G	C	A	A	A
749	A	G	G	A	G	C	A	A	A
749	A	G	G	A	G	C	A	A	A
751	A	G	G	A	G	C	A	A	A
750	A	G	G	A	G	C	A	A	A

FIGURE 18 AA

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Alignment Report of AI-1 alignment, using J. Hein method with Weighted residue weight table.  
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GAGAGCAGGTTGACCATGTGATGATTCCAAAAATCAATCAGGATTTACCA Majority										
9810		9820		9830		9840		9850		
1794	GAGAGCAGGTTGACCATGTGATGATTCCAAAAATCAATCAGGATTTACCA									2603_all.seq
1567	GAGAGCAGGTTGACCATGTGATGATTCCAAAAATCAATCAGGATTTACCA									18rs21_all.seq
1799	GAGAGCAGGTTGACCATGTGATGATTCCAAAAATCAATCAGGATTTACCA									cohl1_all.seq
1799	GAGAGCAGGTTGACCATGTGATGATTCCAAAAATCAATCAGGATTTACCA									cjb111_all.seq
1801	GAGAGCAGGTTGACCATGTGATGATTCCAAAAATCAATCAGGATTTACCA									nem316_all.seq
1800	GAGAGCAGGTTGACCATGTGATGATTCCAAAAATCAATCAGGATTTACCA									a909_all.seq
ATCTACGCTGGTTTCAGAAAGAGGACAATCTGCAACGGGGAGTTGGTCAATCT Majority										
9860		9870		9880		9890		9900		
1844	ATCTACGCTGGTTTCAGAAAGAGGACAATCTGCAACGGGGAGTTGGTCAATCT									2603_all.seq
1617	ATCTACGCTGGTTTCAGAAAGAGGACAATCTGCAACGGGGAGTTGGTCAATCT									18rs21_all.seq
1849	ATCTACGCTGGTTTCAGAAAGAGGACAATCTGCAACGGGGAGTTGGTCAATCT									cohl1_all.seq
1849	ATCTACGCTGGTTTCAGAAAGAGGACAATCTGCAACGGGGAGTTGGTCAATCT									cjb111_all.seq
1851	ATCTACGCTGGTTTCAGAAAGAGGACAATCTGCAACGGGGAGTTGGTCAATCT									nem316_all.seq
1850	ATCTACGCTGGTTTCAGAAAGAGGACAATCTGCAACGGGGAGTTGGTCAATCT									a909_all.seq
AGAAAGGGATAAAGTTTGCCGATTGGAGGGGGCTTCTACACATGCCGCTCTTGA Majority										
9910		9920		9930		9940		9950		
1894	AGAAAGGGATAAAGTTTGCCGATTGGAGGGGGCTTCTACACATGCCGCTCTTGA									2603_all.seq
1667	AGAAAGGGATAAAGTTTGCCGATTGGAGGGGGCTTCTACACATGCCGCTCTTGA									18rs21_all.seq
1899	AGAAAGGGATAAAGTTTGCCGATTGGAGGGGGCTTCTACACATGCCGCTCTTGA									cohl1_all.seq
1899	AGAAAGGGATAAAGTTTGCCGATTGGAGGGGGCTTCTACACATGCCGCTCTTGA									cjb111_all.seq
901	AGAAAGGGATAAAGTTTGCCGATTGGAGGGGGCTTCTACACATGCCGCTCTTGA									nem316_all.seq
900	AGAAAGGGATAAAGTTTGCCGATTGGAGGGGGCTTCTACACATGCCGCTCTTGA									a909_all.seq
GCGGTCAAAGAGGTATGCCAGCTGCTCGGTTGTTTGC GGATTTTGGATAAAG Majority										
9960		9970		9980		9990		10000		
944	GCGGTCAAAGAGGTATGCCAGCTGCTCGGTTGTTTGC GGATTTTGGATAAAG									2603_all.seq
717	GCGGTCAAAGAGGTATGCCAGCTGCTCGGTTGTTTGC GGATTTTGGATAAAG									18rs21_all.seq
949	GCGGTCAAAGAGGTATGCCAGCTGCTCGGTTGTTTGC GGATTTTGGATAAAG									cohl1_all.seq
949	GCGGTCAAAGAGGTATGCCAGCTGCTCGGTTGTTTGC GGATTTTGGATAAAG									cjb111_all.seq
951	GCGGTCAAAGAGGTATGCCAGCTGCTCGGTTGTTTGC GGATTTTGGATAAAG									nem316_all.seq
950	GCGGTCAAAGAGGTATGCCAGCTGCTCGGTTGTTTGC GGATTTTGGATAAAG									a909_all.seq
ATGAAAAAAGGTGATTATTTTTTATGTTACCAATCTGAAAGAAACCTTGGC Majority										
10010		10020		10030		10040		10050		
994	ATGAAAAAAGGTGATTATTTTTTATGTTACCAATCTGAAAGAAACCTTGGC									2603_all.seq
767	ATGAAAAAAGGTGATTATTTTTTATGTTACCAATCTGAAAGAAACCTTGGC									18rs21_all.seq
999	ATGAAAAAAGGTGATTATTTTTTATGTTACCAATCTGAAAGAAACCTTGGC									cohl1_all.seq
999	ATGAAAAAAGGTGATTATTTTTTATGTTACCAATCTGAAAGAAACCTTGGC									cjb111_all.seq
0001	ATGAAAAAAGGTGATTATTTTTTATGTTACCAATCTGAAAGAAACCTTGGC									nem316_all.seq
0000	ATGAAAAAAGGTGATTATTTTTTATGTTACCAATCTGAAAGAAACCTTGGC									a909_all.seq
TTATCAAAGTGGATCGTATCATGCTGATTGAACCTAGCCCAATTGGATGCCCG Majority										
10060		10070		10080		10090		10100		
0044	TTATCAAAGTGGATCGTATCATGCTGATTGAACCTAGCCCAATTGGATGCCCG									2603_all.seq
817	TTATCAAAGTGGATCGTATCATGCTGATTGAACCTAGCCCAATTGGATGCCCG									18rs21_all.seq
0049	TTATCAAAGTGGATCGTATCATGCTGATTGAACCTAGCCCAATTGGATGCCCG									cohl1_all.seq
0049	TTATCAAAGTGGATCGTATCATGCTGATTGAACCTAGCCCAATTGGATGCCCG									cjb111_all.seq
0051	TTATCAAAGTGGATCGTATCATGCTGATTGAACCTAGCCCAATTGGATGCCCG									nem316_all.seq
0050	TTATCAAAGTGGATCGTATCATGCTGATTGAACCTAGCCCAATTGGATGCCCG									a909_all.seq
TGAGCATTGGAAGAGGATAAAGATTATGTTACCCCTTCTGACCTGTACACCT Majority										
10110		10120		10130		10140		10150		
0094	TGAGCATTGGAAGAGGATAAAGATTATGTTACCCCTTCTGACCTGTACACCT									2603_all.seq
867	TGAGCATTGGAAGAGGATAAAGATTATGTTACCCCTTCTGACCTGTACACCT									18rs21_all.seq
0099	TGAGCATTGGAAGAGGATAAAGATTATGTTACCCCTTCTGACCTGTACACCT									cohl1_all.seq
0099	TGAGCATTGGAAGAGGATAAAGATTATGTTACCCCTTCTGACCTGTACACCT									cjb111_all.seq
0101	TGAGCATTGGAAGAGGATAAAGATTATGTTACCCCTTCTGACCTGTACACCT									nem316_all.seq
0100	TGAGCATTGGAAGAGGATAAAGATTATGTTACCCCTTCTGACCTGTACACCT									a909_all.seq

FIGURE 18 AB



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T A T A T G G G C T C T T T G T C A A C T G T A A T G G G T G A C T T A T C A T T A A C A A C G A G Majority									
10160		10170		10180		10190		10200	
10144	T A T A T G G G C T C T T T G T C A A C T G T A A T G G G T G A C T T A T C A T T A A C A A C G A G 2603_all.seq								
9917	T A T A T G G G C T C T T T G T C A A C T G T A A T G G G T G A C T T A T C A T T A A C A A C G A G 18rs21_all.seq								
10149	T A T A T G G G C T C T T T G T C A A C T G T A A T G G G T G A C T T A T C A T T A A C A A C G A G cohl_all.seq								
10149	T A T A T G G G C T C T T T G T C A A C T G T A A T G G G T G A C T T A T C A T T A A C A A C G A G cjb111_all.seq								
10151	T A T A T G G G C T C T T T G T C A A C T G T A A T G G G T G A C T T A T C A T T A A C A A C G A G nem316_all.seq								
10150	T A T A T G G G C T C T T T G T C A A C T G T A A T G G G T G A C T T A T C A T T A A C A A C G A G a909_all.seq								
A G A G A A T C A G C T T G G T T C T C T C T T T T T G G A T G T T C A A A G C G A T G A G A A Majority									
10210		10220		10230		10240		10250	
10194	A G A G A A T C A G C T T G G T T C T C T C T C T T T T T G G A T G T T C A A A G C G A T G A G A A 2603_all.seq								
9967	A G A G A A T C A G C T T G G T T C T C T C T C T T T T T G G A T G T T C A A A G C G A T G A G A A 18rs21_all.seq								
10199	A G A G A A T C A G C T T G G T T C T C T C T C T T T T T G G A T G T T C A A A G C G A T G A G A A cohl_all.seq								
10199	A G A G A A T C A G C T T G G T T C T C T C T C T T T T T G G A T G T T C A A A G C G A T G A G A A cjb111_all.seq								
10201	A G A G A A T C A G C T T G G T T C T C T C T C T T T T T G G A T G T T C A A A G C G A T G A G A A nem316_all.seq								
10200	A G A G A A T C A G C T T G G T T C T C T C T C T T T T T G G A T G T T C A A A G C G A T G A G A A a909_all.seq								
T T T T G C T T T T A A A A T T T T T A A A G T T G C G A A A G C C A A A G G C T T G T C G C T T G Majority									
10260		10270		10280		10290		10300	
10244	T T T T G C T T T T A A A A T T T T T A A A G T T G C G A A A G C C A A A G G C T T G T C G C T T G 2603_all.seq								
10017	T T T T G C T T T T A A A A T T T T T A A A G T T G C G A A A G C C A A A G G C T T G T C G C T T G 18rs21_all.seq								
10249	T T T T G C T T T T A A A A T T T T T A A A G T T G C G A A A G C C A A A G G C T T G T C G C T T G cohl_all.seq								
10249	T T T T G C T T T T A A A A T T T T T A A A G T T G C G A A A G C C A A A G G C T T G T C G C T T G cjb111_all.seq								
10251	T T T T G C T T T T A A A A T T T T T A A A G T T G C G A A A G C C A A A G G C T T G T C G C T T G nem316_all.seq								
10250	T T T T G C T T T T A A A A T T T T T A A A G T T G C G A A A G C C A A A G G C T T G T C G C T T G a909_all.seq								
A T G T C T T T G A T A A G C T T G T T G G T C G C T T C T A A T T T G G C A T T G G A A A G G G A Majority									
10310		10320		10330		10340		10350	
10294	A T G T C T T T G A T A A G C T T G T T G G T C G C T T C T A A T T T G G C A T T G G A A A G G G A 2603_all.seq								
10067	A T G T C T T T G A T A A G C T T G T T G G T C G C T T C T A A T T T G G C A T T G G A A A G G G A 18rs21_all.seq								
10299	A T G T C T T T G A T A A G C T T G T T G G T C G C T T C T A A T T T G G C A T T G G A A A G G G A cohl_all.seq								
10299	A T G T C T T T G A T A A G C T T G T T G G T C G C T T C T A A T T T G G C A T T G G A A A G G G A cjb111_all.seq								
10301	A T G T C T T T G A T A A G C T T G T T G G T C G C T T C T A A T T T G G C A T T G G A A A G G G A nem316_all.seq								
10300	A T G T C T T T G A T A A G C T T G T T G G T C G C T T C T A A T T T G G C A T T G G A A A G G G A a909_all.seq								
T A G C T G A G G G C G T T T G T G A T G T A A G T T T T G T A C T T C A T G A A G G T C C T A A A Majority									
10360		10370		10380		10390		10400	
10344	T A G C T G A G G G C G T T T G T G A T G T A A G T T T T G T A C T T C A T G A A G G T C C T A A A 2603_all.seq								
10117	T A G C T G A G G G C G T T T G T G A T G T A A G T T T T G T A C T T C A T G A A G G T C C T A A A 18rs21_all.seq								
10349	T A G C T G A G G G C G T T T G T G A T G T A A G T T T T G T A C T T C A T G A A G G T C C T A A A cohl_all.seq								
10349	T A G C T G A G G G C G T T T G T G A T G T A A G T T T T G T A C T T C A T G A A G G T C C T A A A cjb111_all.seq								
10351	T A G C T G A G G G C G T T T G T G A T G T A A G T T T T G T A C T T C A T G A A G G T C C T A A A nem316_all.seq								
10350	T A G C T G A G G G C G T T T G T G A T G T A A G T T T T G T A C T T C A T G A A G G T C C T A A A a909_all.seq								
G A C C A T T T T A A A A G C A G A A T T G C C A G T T G T T A C G T T A T G A T C T A T G A A G C Majority									
10410		10420		10430		10440		10450	
0394	G A C C A T T T T A A A A G C A G A A T T G C C A G T T G T T A C G T T A T G A T C T A T G A A G C 2603_all.seq								
0167	G A C C A T T T T A A A A G C A G A A T T G C C A G T T G T T A C G T T A T G A T C T A T G A A G C 18rs21_all.seq								
0399	G A C C A T T T T A A A A G C A G A A T T G C C A G T T G T T A C G T T A T G A T C T A T G A A G C cohl_all.seq								
0399	G A C C A T T T T A A A A G C A G A A T T G C C A G T T G T T A C G T T A T G A T C T A T G A A G C cjb111_all.seq								
0401	G A C C A T T T T A A A A G C A G A A T T G C C A G T T G T T A C G T T A T G A T C T A T G A A G C nem316_all.seq								
0400	G A C C A T T T T A A A A G C A G A A T T G C C A G T T G T T A C G T T A T G A T C T A T G A A G C a909_all.seq								
T A A A G A A T T C A G T T G T T C G C T T T T C T T A G A A A T G A A A G A G T A T G A G C T G A Majority									
10460		10470		10480		10490		10500	
0444	T A A A G A A T T C A G T T G T T C G C T T T T C T T A G A A A T G A A A G A G T A T G A G C T G A 2603_all.seq								
0217	T A A A G A A T T C A G T T G T T C G C T T T T C T T A G A A A T G A A A G A G T A T G A G C T G A 18rs21_all.seq								
0449	T A A A G A A T T C A G T T G T T C G C T T T T C T T A G A A A T G A A A G A G T A T G A G C T G A cohl_all.seq								
0449	T A A A G A A T T C A G T T G T T C G C T T T T C T T A G A A A T G A A A G A G T A T G A G C T G A cjb111_all.seq								
0451	T A A A G A A T T C A G T T G T T C G C T T T T C T T A G A A A T G A A A G A G T A T G A G C T G A nem316_all.seq								
0450	T A A A G A A T T C A G T T G T T C G C T T T T C T T A G A A A T G A A A G A G T A T G A G C T G A a909_all.seq								

FIGURE 18 AC

T A A A G G T C G T T A T A A T A G C G G A G C T C A T C T G A G A A A A C T T T T A T A C C T C A Majority									
10510		10520		10530		10540		10550	
10494	T A A A G G T C G T T A T A A T A G C G G A G C T C A T C T G A G A A A A C T T T T A T A C C T C A	2603_all.seq							
10267	T A A A G G T C G T T A T A A T A G C G G A G C T C A T C T G A G A A A A C T T T T A T A C C T C A	18rs21_all.seq							
10499	T A A A G G T C G T T A T A A T A G C G G A G C T C A T C T G A G A A A A C T T T T A T A C C T C A	cohl_all.seq							
10499	T A A A G G T C G T T A T A A T A G C G G A G C T C A T C T G A G A A A A C T T T T A T A C C T C A	cjb111_all.seq							
10501	T A A A G G T C G T T A T A A T A G C G G A G C T C A T C T G A G A A A A C T T T T A T A C C T C A	nem316_all.seq							
10500	T A A A G G T C G T T A T A A T A G C G G A G C T C A T C T G A G A A A A C T T T T A T A C C T C A	a909_all.seq							
A A G T C A G T C T A G C T T T G A T A T C A T G A A G C C A T T A G G A G T T A T T C C T T A T C Majority									
10560		10570		10580		10590		10600	
10544	A A G T C A G T C T A G C T T T G A T A T C A T G A A G C C A T T A G G A G T T A T T C C T T A T C	2603_all.seq							
10317	A A G T C A G T C T A G C T T T G A T A T C A T G A A G C C A T T A G G A G T T A T T C C T T A T C	18rs21_all.seq							
10549	A A G T C A G T C T A G C T T T G A T A T C A T G A A G C C A T T A G G A G T T A T T C C T T A T C	cohl_all.seq							
10549	A A G T C A G T C T A G C T T T G A T A T C A T G A A G C C A T T A G G A G T T A T T C C T T A T C	cjb111_all.seq							
10551	A A G T C A G T C T A G C T T T G A T A T C A T G A A G C C A T T A G G A G T T A T T C C T T A T C	nem316_all.seq							
10550	A A G T C A G T C T A G C T T T G A T A T C A T G A A G C C A T T A G G A G T T A T T C C T T A T C	a909_all.seq							
T T T T A G T G C C G C G C G A T C C A T A T A G T G A T A G A T C G A G A T A T T T A G A T C C A Majority									
10610		10620		10630		10640		10650	
10594	T T T T A G T G C C G C G C G A T C C A T A T A G T G A T A G A T C G A G A T A T T T A G A T C C A	2603_all.seq							
10367	T T T T A G T G C C G C G C G A T C C A T A T A G T G A T A G A T C G A G A T A T T T A G A T C C A	18rs21_all.seq							
10599	T T T T A G T G C C G C G C G A T C C A T A T A G T G A T A G A T C G A G A T A T T T A G A T C C A	cohl_all.seq							
10599	T T T T A G T G C C G C G C G A T C C A T A T A G T G A T A G A T C G A G A T A T T T A G A T C C A	cjb111_all.seq							
10601	T T T T A G T G C C G C G C G A T C C A T A T A G T G A T A G A T C G A G A T A T T T A G A T C C A	nem316_all.seq							
10600	T T T T A G T G C C G C G C G A T C C A T A T A G T G A T A G A T C G A G A T A T T T A G A T C C A	a909_all.seq							
A A A G T T C T A T C A T C C T C T T T T G G C G C C T T T T T T C C A G C A G A T A A T A T T A A Majority									
10660		10670		10680		10690		10700	
10644	A A A G T T C T A T C A T C C T C T T T T G G C G C C T T T T T T C C A G C A G A T A A T A T T A A	2603_all.seq							
10417	A A A G T T C T A T C A T C C T C T T T T G G C G C C T T T T T T C C A G C A G A T A A T A T T A A	18rs21_all.seq							
10649	A A A G T T C T A T C A T C C T C T T T T G G C G C C T T T T T T C C A G C A G A T A A T A T T A A	cohl_all.seq							
10649	A A A G T T C T A T C A T C C T C T T T T G G C G C C T T T T T T C C A G C A G A T A A T A T T A A	cjb111_all.seq							
10651	A A A G T T C T A T C A T C C T C T T T T G G C G C C T T T T T T C C A G C A G A T A A T A T T A A	nem316_all.seq							
10650	A A A G T T C T A T C A T C C T C T T T T G G C G C C T T T T T T C C A G C A G A T A A T A T T A A	a909_all.seq							
G G T A G C T T G G T C T A A C A A C T C C A G C A G T T T A T T T A C A C C A C C T A T T A A T G Majority									
10710		10720		10730		10740		10750	
0694	G G T A G C T T G G T C T A A C A A C T C C A G C A C T T T A T T T A C A C C A C C T A T T A A T G	2603_all.seq							
0467	G G T A G C T T G G T C T A A C A A C T C C A G C A C T T T A T T T A C A C C A C C T A T T A A T G	18rs21_all.seq							
0699	G G T A G C T T G G T C T A A C A A C T C C A G C A C T T T A T T T A C A C C A C C T A T T A A T G	cohl_all.seq							
0699	G G T A G C T T G G T C T A A C A A C T C C A G C A C T T T A T T T A C A C C A C C T A T T A A T G	cjb111_all.seq							
0701	G G T A G C T T G G T C T A A C A A C T C C A G C A C T T T A T T T A C A C C A C C T A T T A A T G	nem316_all.seq							
0700	G G T A G C T T G G T C T A A C A A C T C C A G C A C T T T A T T T A C A C C A C C T A T T A A T G	a909_all.seq							
C A A A C T A C A C C A C T C A G A T T C A A G C T A T T G G G A C A A C G A T T A A G T C A C A A Majority									
10760		10770		10780		10790		10800	
0744	C A A A C T A C A C C A C T C A G A T T C A A G C T A T T G G G A C A A C G A T T A A G T C A C A A	2603_all.seq							
0517	C A A A C T A C A C C A C T C A G A T T C A A G C T A T T G G G A C A A C G A T T A A G T C A C A A	18rs21_all.seq							
0749	C A A A C T A C A C C A C T C A G A T T C A A G C T A T T G G G A C A A C G A T T A A G T C A C A A	cohl_all.seq							
0749	C A A A C T A C A C C A C T C A G A T T C A A G C T A T T G G G A C A A C G A T T A A G T C A C A A	cjb111_all.seq							
0751	C A A A C T A C A C C A C T C A G A T T C A A G C T A T T G G G A C A A C G A T T A A G T C A C A A	nem316_all.seq							
0750	C A A A C T A C A C C A C T C A G A T T C A A G C T A T T G G G A C A A C G A T T A A G T C A C A A	a909_all.seq							
A T T C C G G A A T C G A T T T T G A C G G T T A C G G A T A A A A A G A G C A G G A A G T T C A G Majority									
10810		10820		10830		10840		10850	
0794	A T T C C G G A A T C G A T T T T G A C G G T T A C G G A T A A A A A G A G C A G G A A G T T C A G	2603_all.seq							
0567	A T T C C G G A A T C G A T T T T G A C G G T T A C G G A T A A A A A G A G C A G G A A G T T C A G	18rs21_all.seq							
0799	A T T C C G G A A T C G A T T T T G A C G G T T A C G G A T A A A A A G A G C A G G A A G T T C A G	cohl_all.seq							
0799	A T T C C G G A A T C G A T T T T G A C G G T T A C G G A T A A A A A G A G C A G G A A G T T C A G	cjb111_all.seq							
0801	A T T C C G G A A T C G A T T T T G A C G G T T A C G G A T A A A A A G A G C A G G A A G T T C A G	nem316_all.seq							
0800	A T T C C G G A A T C G A T T T T G A C G G T T A C G G A T A A A A A G A G C A G G A A G T T C A G	a909_all.seq							

FIGURE 18 AD



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C A T T A A C A A G A T T G A C G A A G C T A A A G A A G G C T T A G T A G G T G C G A C C T T C A Majority									
10860		10870		10880		10890		10900	
10844	C	A	T	T	A	A	C	A	A
10617	C	A	T	T	A	A	C	A	A
10849	C	A	T	T	A	A	C	A	A
10849	C	A	T	T	A	A	C	A	A
10851	C	A	T	T	A	A	C	A	A
10850	C	A	T	T	A	A	C	A	A
C C T T G T C T A A A C G C A C A A C A G T A G C G G C A G A T C A T C A A G T A C A A G G A G A T Majority									
10910		10920		10930		10940		10950	
10894	C	C	T	T	G	T	C	T	A
10667	C	C	T	T	G	T	C	T	A
10899	C	C	T	T	G	T	C	T	A
10899	C	C	T	T	G	T	C	T	A
10901	C	C	T	T	G	T	C	T	A
10900	C	C	T	T	G	T	C	T	A
T T C A T T C C T G T C A G C A A A G A G A C G A C A G T C G G T C G G A C A A C C C T T A C C T T Majority									
10960		10970		10980		10990		11000	
10944	T	T	C	A	T	T	C	C	T
10717	T	T	C	A	T	T	C	C	T
10949	T	T	C	A	T	T	C	C	T
10949	T	T	C	A	T	T	C	C	T
10951	T	T	C	A	T	T	C	C	T
10950	T	T	C	A	T	T	C	C	T
T G A C A A C C T T A A A C C T G G A T T T T A T G A C C T T A A A G A A A C G A A A G C G C C G A Majority									
11010		11020		11030		11040		11050	
0994	T	G	A	C	A	A	C	C	T
0767	T	G	A	C	A	A	C	C	T
0999	T	G	A	C	A	A	C	C	T
0999	T	G	A	C	A	A	C	C	T
1001	T	G	A	C	A	A	C	C	T
1000	T	G	A	C	A	A	C	C	T
A T G C T T A C G T A C T T G A T C C T A A G A C T T A T G T T G T G G T C G T T C A A A A T T C A Majority									
11060		11070		11080		11090		11100	
1044	A	T	G	C	T	T	A	C	G
10817	A	T	G	C	T	T	A	C	G
1049	A	T	G	C	T	T	A	C	G
1049	A	T	G	C	T	T	A	C	G
1051	A	T	G	C	T	T	A	C	G
1050	A	T	G	C	T	T	A	C	G
G G A A A A A C G A C A A T T G T G G A T G A A G C T A A C T T C A A A G A G G C T G A T T A C C C Majority									
11110		11120		11130		11140		11150	
1094	G	G	A	A	A	A	A	C	G
1067	G	G	A	A	A	A	A	C	G
1099	G	G	A	A	A	A	A	C	G
1099	G	G	A	A	A	A	A	C	G
1101	G	G	A	A	A	A	A	C	G
1100	G	G	A	A	A	A	A	C	G
A A T G G C T G A T A A T A C C A G C C A T G T G G A G T G C G T A G C G T T G C T A C A A C G A A Majority									
11160		11170		11180		11190		11200	
144	A	A	T	G	G	C	T	G	A
1917	A	A	T	G	G	C	T	G	A
149	A	A	T	G	G	C	T	G	A
149	A	A	T	G	G	C	T	G	A
151	A	A	T	G	G	C	T	G	A
150	A	A	T	G	G	C	T	G	A

FIGURE 18 AE

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G C A A A G G G T A A A A A T C C T T T A T T T T A A G C A C T T T T T C A A G C A T T T T G T C T Majority										
11210		11220		11230		11240		11250		
11194	G C A A A G G G T A A A A A T C C T T T A T T T T A A G C A C T T T T T C A A G C A T T T T G T C T									2603_all.seq
10967	G C A A A G G G T A A A A A T C C T T T A T T T T A A G C A C T T T T T C A A G C A T T T T G T C T									18rs21_all.seq
11199	G C A A A G G G T A A A A A T C C T T T A T T T T A A G C A C T T T T T C A A G C A T T T T G T C T									cohl_all.seq
11199	G C A A A G G G T A A A A A T C C T T T A T T T T A A G C A C T T T T T C A A G C A T T T T G T C T									cjb111_all.seq
11201	G C A A A G G G T A A A A A T C C T T T A T T T T A A G C A C T T T T T C A A G C A T T T T G T C T									nem316_all.seq
11200	G C A A A G G G T A A A A A T C C T T T A T T T T A A G C A C T T T T T C A A G C A T T T T G T C T									a909_all.seq
T T A T T G A A A A G A G T G A T T T T A A C A T A A A A A A G G T A T T A A A A A A C A T A T T G Majority										
11260		11270		11280		11290		11300		
11244	T T A T T G A A A A G A G T G A T T T T A A C A T A A A A A A G G T A T T A A A A A A C A T A T T G									2603_all.seq
11017	T T A T T G A A A A G A G T G A T T T T A A C A T A A A A A A G G T A T T A A A A A A C A T A T T G									18rs21_all.seq
11249	T T A T T G A A A A G A G T G A T T T T A A C A T A A A A A A G G T A T T A A A A A A C A T A T T G									cohl_all.seq
11249	T T A T T G A A A A G A G T G A T T T T A A C A T A A A A A A G G T A T T A A A A A A C A T A T T G									cjb111_all.seq
11251	T T A T T G A A A A G A G T G A T T T T A A C A T A A A A A A G G T A T T A A A A A A C A T A T T G									nem316_all.seq
11250	T T A T T G A A A A G A G T G A T T T T A A C A T A A A A A A G G T A T T A A A A A A C A T A T T G									a909_all.seq
A C G T G A C C G T T T G T T T T G A A G T G G C T T G C G T A G A C A A A A A A A T A G A T A C G Majority										
11310		11320		11330		11340		11350		
11294	A C G T G A C C G T T T G T T T T G A A G T G G C T T G C G T A G A C A A A A A A A T A G A T A C G									2603_all.seq
11067	A C G T G A C C G T T T G T T T T G A A G T G G C T T G C G T A G A C A A A A A A A T A G A T A C G									18rs21_all.seq
11299	A C G T G A C C G T T T G T T T T G A A G T G G C T T G C G T A G A C A A A A A A A T A G A T A C G									cohl_all.seq
11299	A C G T G A C C G T T T G T T T T G A A G T G G C T T G C G T A G A C A A A A A A A T A G A T A C G									cjb111_all.seq
11301	A C G T G A C C G T T T G T T T T G A A G T G G C T T G C G T A G A C A A A A A A A T A G A T A C G									nem316_all.seq
11300	A C G T G A C C G T T T G T T T T G A A G T G G C T T G C G T A G A C A A A A A A A T A G A T A C G									a909_all.seq
T C A G A T A A A T T T C T G G C A T T A C G A G A A C A T T T T T A G A G T G T T C T C T T T T T Majority										
11360		11370		11380		11390		11400		
11344	T C A G A T A A A T T T C T G G C A T T A C G A G A A C A T T T T T A G A G T G T T C T C T T T T T									2603_all.seq
11117	T C A G A T A A A T T T C T G G C A T T A C G A G A A C A T T T T T A G A G T G T T C T C T T T T T									18rs21_all.seq
11349	T C A G A T A A A T T T C T G G C A T T A C G A G A A C A T T T T T A G A G T G T T C T C T T T T T									cohl_all.seq
11349	T C A G A T A A A T T T C T G G C A T T A C G A G A A C A T T T T T A G A G T G T T C T C T T T T T									cjb111_all.seq
11351	T C A G A T A A A T T T C T G G C A T T A C G A G A A C A T T T T T A G A G T G T T C T C T T T T T									nem316_all.seq
11350	T C A G A T A A A T T T C T G G C A T T A C G A G A A C A T T T T T A G A G T G T T C T C T T T T T									a909_all.seq
T T A G T T T A C G G A G G A A A A A T A T A T A T G G A A A A A C A G G A T T C A C G A G T T C T Majority										
11410		11420		11430		11440		11450		
11394	T T A G T T T A C G G A G G A A A A A T A T A T A T G G A A A A A C A G G A T T C A C G A G T T C T									2603_all.seq
11167	T T A G T T T A C G G A G G A A A A A T A T A T A T G G A A A A A C A G G A T T C A C G A G T T C T									18rs21_all.seq
11399	T T A G T T T A C G G A G G A A A A A T A T A T A T G G A A A A A C A G G A T T C A C G A G T T C T									cohl_all.seq
11399	T T A G T T T A C G G A G G A A A A A T A T A T A T G G A A A A A C A G G A T T C A C G A G T T C T									cjb111_all.seq
1401	T T A G T T T A C G G A G G A A A A A T A T A T A T G G A A A A A C A G G A T T C A C G A G T T C T									nem316_all.seq
1400	T T A G T T T A C G G A G G A A A A A T A T A T A T G G A A A A A C A G G A T T C A C G A G T T C T									a909_all.seq
C A T C C A T T G G G A G G G G A A T T C T G G G G A C A A G C T C A T T G A A C A C C A A A C C A Majority										
11460		11470		11480		11490		11500		
1444	C A T C C A T T G G G A G G G G A A T T C T G G G G A C A A G C T C A T T G A A C A C C A A A C C A									2603_all.seq
1217	C A T C C A T T G G G A G G G G A A T T C T G G G G A C A A G C T C A T T G A A C A C C A A A C C A									18rs21_all.seq
1449	C A T C C A T T G G G A G G G G A A T T C T G G G G A C A A G C T C A T T G A A C A C C A A A C C A									cohl_all.seq
1449	C A T C C A T T G G G A G G G G A A T T C T G G G G A C A A G C T C A T T G A A C A C C A A A C C A									cjb111_all.seq
1451	C A T C C A T T G G G A G G G G A A T T C T G G G G A C A A G C T C A T T G A A C A C C A A A C C A									nem316_all.seq
1450	C A T C C A T T G G G A G G G G A A T T C T G G G G A C A A G C T C A T T G A A C A C C A A A C C A									a909_all.seq
G C G C A A C G G G G T G G T A C T A C C A A G T C G A T C G T A G C T T T A G T C A A C C A A A A Majority										
11510		11520		11530		11540		11550		
1494	G C G C A A C G G G G T G G T A C T A C C A A G T C G A T C G T A G C T T T A G T C A A C C A A A A									2603_all.seq
1267	G C G C A A C G G G G T G G T A C T A C C A A G T C G A T C G T A G C T T T A G T C A A C C A A A A									18rs21_all.seq
1499	G C G C A A C G G G G T G G T A C T A C C A A G T C G A T C G T A G C T T T A G T C A A C C A A A A									cohl_all.seq
1499	G C G C A A C G G G G T G G T A C T A C C A A G T C G A T C G T A G C T T T A G T C A A C C A A A A									cjb111_all.seq
1501	G C G C A A C G G G G T G G T A C T A C C A A G T C G A T C G T A G C T T T A G T C A A C C A A A A									nem316_all.seq
1500	G C G C A A C G G G G T G G T A C T A C C A A G T C G A T C G T A G C T T T A G T C A A C C A A A A									a909_all.seq

FIGURE 18 AF



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G G G G A A C C G C C C A G A A T G A T C G A A A G C C A C T A G A A A G T G T C C G T A A T G A T Majority									
11560		11570		11580		11590		11600	
11544	G	G	G	G	A	A	C	C	G
11317	G	G	G	G	A	A	C	C	G
11549	G	G	G	G	A	A	C	C	G
11549	G	G	G	G	A	A	C	C	G
11551	G	G	G	G	A	A	C	C	G
11550	G	G	G	G	A	A	C	C	G
T C G A T T T C G G G C G G T G A T G A T G T C A T G G G T T A T G C T T A T A G C A A A T G T A C Majority									
11610		11620		11630		11640		11650	
11594	T	C	G	A	T	T	C	G	G
11367	T	C	G	A	T	T	C	G	G
11599	T	C	G	A	T	T	C	G	G
11599	T	C	G	A	T	T	C	G	G
11601	T	C	G	A	T	T	C	G	G
11600	T	C	G	A	T	T	C	G	G
T T G G G G A G T T G C G G C A C G A A T T A A T C A G T G G G A C T G A A A C T C A A A G G T T G Majority									
11660		11670		11680		11690		11700	
1644	T	T	G	G	G	A	G	T	T
1417	T	T	G	G	G	A	G	T	T
1649	T	T	G	G	G	A	G	T	T
1649	T	T	G	G	G	A	G	T	T
1651	T	T	G	G	G	A	G	T	T
1650	T	T	G	G	G	A	G	T	T
A A A T G G T G A G A A G A T T A C C A T T A C C A G T T C A A T G G G A A A T G G T C A G G A T T Majority									
11710		11720		11730		11740		11750	
1694	A	A	A	T	G	G	T	G	A
1467	A	A	A	T	G	G	T	G	A
1699	A	A	A	T	G	G	T	G	A
1699	A	A	A	T	G	G	T	G	A
1701	A	A	A	T	G	G	T	G	A
1700	A	A	A	T	G	G	T	G	A
G G G T T G G A A C A G C C G A A A G A C T A G A T G G T G A A A C T G A T A C A G T T C C A A A A Majority									
11760		11770		11780		11790		11800	
1744	G	G	G	T	T	G	G	A	A
1517	G	G	G	T	T	G	G	A	A
1749	G	G	G	T	T	G	G	A	A
1749	G	G	G	T	T	G	G	A	A
1751	G	G	G	T	T	G	G	A	A
1750	G	G	G	T	T	G	G	A	A
G A A G G T A C T A T T C T C T C T T T T A G G A A A G T A G T T A T G C T T C G T A T A T A G G Majority									
11810		11820		11830		11840		11850	
794	G	A	A	G	G	T	A	C	T
567	G	A	A	G	G	T	A	C	T
799	G	A	A	G	G	T	A	C	T
799	G	A	A	G	G	T	A	C	T
801	G	A	A	G	G	T	A	C	T
800	G	A	A	G	G	T	A	C	T
C T A C G G A A C T A T A T C T T T C G T C A C A T T A C A T C T A C A G A T A G T A C C A T G A A Majority									
11860		11870		11880		11890		11900	
844	C	T	A	C	G	G	A	A	C
617	C	T	A	C	G	G	A	A	C
849	C	T	A	C	G	G	A	A	C
849	C	T	A	C	G	G	A	A	C
851	C	T	A	C	G	G	A	A	C
850	C	T	A	C	G	G	A	A	C

FIGURE 18 AG

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Alignment Report of AI-1 alignment, using J. Hein method with Weighted residue weight table.  
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TTTTGCTTATATGACCAAGTAAAGTGAGGATATACTAACAAATGAAATAT										Majority
11910		11920		11930		11940		11950		
11894	TTTTGCTTATATGACCAAGTAAAGTGAGGATATACTAACAAATGAAATAT									2603_all.seq
11667	TTTTGCTTATATGACCAAGTAAAGTGAGGATATACTAACAAATGAAATAT									18rs21_all.seq
11899	TTTTGCTTATATGACCAAGTAAAGTGAGGATATACTAACAAATGAAATAT									cohl_all.seq
11899	TTTTGCTTATATGACCAAGTAAAGTGAGGATATACTAACAAATGAAATAT									cjb111_all.seq
11901	TTTTGCTTATATGACCAAGTAAAGTGAGGATATACTAACAAATGAAATAT									nem316_all.seq
11900	TTTTGCTTATATGACCAAGTAAAGTGAGGATATACTAACAAATGAAATAT									a909_all.seq
TTATTATCGTATTTTGTCCATTTTATCGAAAAGTTTGCATATTATCATTAT										Majority
11960		11970		11980		11990		12000		
11944	TTATTATCGTATTTTGTCCATTTTATCGAAAAGTTTGCATATTATCATTAT									2603_all.seq
11717	TTATTATCGTATTTTGTCCATTTTATCGAAAAGTTTGCATATTATCATTAT									18rs21_all.seq
11949	TTATTATCGTATTTTGTCCATTTTATCGAAAAGTTTGCATATTATCATTAT									cohl_all.seq
11949	TTATTATCGTATTTTGTCCATTTTATCGAAAAGTTTGCATATTATCATTAT									cjb111_all.seq
11951	TTATTATCGTATTTTGTCCATTTTATCGAAAAGTTTGCATATTATCATTAT									nem316_all.seq
11950	TTATTATCGTATTTTGTCCATTTTATCGAAAAGTTTGCATATTATCATTAT									a909_all.seq
GTTTGATAAAGATGCAAATATAATGATAGTAGGAGCTAAATATGGATATTTT										Majority
12010		12020		12030		12040		12050		
11994	GTTTGATAAAGATGCAAATATAATGATAGTAGGAGCTAAATATGGATATTTT									2603_all.seq
11767	GTTTGATAAAGATGCAAATATAATGATAGTAGGAGCTAAATATGGATATTTT									18rs21_all.seq
11999	GTTTGATAAAGATGCAAATATAATGATAGTAGGAGCTAAATATGGATATTTT									cohl_all.seq
11999	GTTTGATAAAGATGCAAATATAATGATAGTAGGAGCTAAATATGGATATTTT									cjb111_all.seq
12001	GTTTGATAAAGATGCAAATATAATGATAGTAGGAGCTAAATATGGATATTTT									nem316_all.seq
12000	GTTTGATAAAGATGCAAATATAATGATAGTAGGAGCTAAATATGGATATTTT									a909_all.seq
AAAAAATCAAGAGTATCCTAAGTGCTTTTCCATTTTGA AATTCA AATATAG										Majority
12060		12070		12080		12090		12100		
12044	AAAAAATCAAGAGTATCCTAAGTGCTTTTCCATTTTGA AATTCA AATATAG									2603_all.seq
11817	AAAAAATCAAGAGTATCCTAAGTGCTTTTCCATTTTGA AATTCA AATATAG									18rs21_all.seq
12049	AAAAAATCAAGAGTATCCTAAGTGCTTTTCCATTTTGA AATTCA AATATAG									cohl_all.seq
12049	AAAAAATCAAGAGTATCCTAAGTGCTTTTCCATTTTGA AATTCA AATATAG									cjb111_all.seq
12051	AAAAAATCAAGAGTATCCTAAGTGCTTTTCCATTTTGA AATTCA AATATAG									nem316_all.seq
12050	AAAAAATCAAGAGTATCCTAAGTGCTTTTCCATTTTGA AATTCA AATATAG									a909_all.seq
CTAATAGTTCTAGAACTTCTAATTGTTTTTTCGTCGACGATATGAATTTTTC										Majority
12110		12120		12130		12140		12150		
2094	CTAATAGTTCTAGAACTTCTAATTGTTTTTTCGTCGACGATATGAATTTTTC									2603_all.seq
1867	CTAATAGTTCTAGAACTTCTAATTGTTTTTTCGTCGACGATATGAATTTTTC									18rs21_all.seq
2099	CTAATAGTTCTAGAACTTCTAATTGTTTTTTCGTCGACGATATGAATTTTTC									cohl_all.seq
2099	CTAATAGTTCTAGAACTTCTAATTGTTTTTTCGTCGACGATATGAATTTTTC									cjb111_all.seq
2101	CTAATAGTTCTAGAACTTCTAATTGTTTTTTCGTCGACGATATGAATTTTTC									nem316_all.seq
2100	CTAATAGTTCTAGAACTTCTAATTGTTTTTTCGTCGACGATATGAATTTTTC									a909_all.seq
AATCTTAACCTGTTAGGATTCCACCTCCCTTTTGGTTAAAGAAAAAAGGTC A										Majority
12160		12170		12180		12190		12200		
2144	AATCTTAACCTGTTAGGATTCCACCTCCCTTTTGGTTAAAGAAAAAAGGTC A									2603_all.seq
1917	AATCTTAACCTGTTAGGATTCCACCTCCCTTTTGGTTAAAGAAAAAAGGTC A									18rs21_all.seq
2149	AATCTTAACCTGTTAGGATTCCACCTCCCTTTTGGTTAAAGAAAAAAGGTC A									cohl_all.seq
2149	AATCTTAACCTGTTAGGATTCCACCTCCCTTTTGGTTAAAGAAAAAAGGTC A									cjb111_all.seq
2151	AATCTTAACCTGTTAGGATTCCACCTCCCTTTTGGTTAAAGAAAAAAGGTC A									nem316_all.seq
2150	AATCTTAACCTGTTAGGATTCCACCTCCCTTTTGGTTAAAGAAAAAAGGTC A									a909_all.seq
GGTCGTTTATAGATAA AACTTTGTCAAACAAGCTCAAGCTATCTAAAAAATAGTT										Majority
12210		12220		12230		12240		12250		
2194	GGTCGTTTATAGATAA AACTTTGTCAAACAAGCTCAAGCTATCTAAAAAATAGTT									2603_all.seq
1967	GGTCGTTTATAGATAA AACTTTGTCAAACAAGCTCAAGCTATCTAAAAAATAGTT									18rs21_all.seq
2199	GGTCGTTTATAGATAA AACTTTGTCAAACAAGCTCAAGCTATCTAAAAAATAGTT									cohl_all.seq
2199	GGTCGTTTATAGATAA AACTTTGTCAAACAAGCTCAAGCTATCTAAAAAATAGTT									cjb111_all.seq
2201	GGTCGTTTATAGATAA AACTTTGTCAAACAAGCTCAAGCTATCTAAAAAATAGTT									nem316_all.seq
2200	GGTCGTTTATAGATAA AACTTTGTCAAACAAGCTCAAGCTATCTAAAAAATAGTT									a909_all.seq

FIGURE 18 AH



T G A A A T G G G C A T T A C T C T A G T T T T T A A T A A G C T A T C T G A T G A G C A G A A G G Majority									
12260		12270		12280		12290		12300	
12244	T	G	A	A	A	T	G	G	G
12017	T	G	A	A	A	T	G	G	G
12249	T	G	A	A	A	T	G	G	G
12249	T	G	A	A	A	T	G	G	G
12251	T	G	A	A	A	T	G	G	G
12250	T	G	A	A	A	T	G	G	G
A G A A G T T A A T G C A T G T T G G G A A G T C T T A T T T T G A C T A T C A A G A A A A T G C T Majority									
12310		12320		12330		12340		12350	
12294	A	G	A	A	G	T	T	A	A
12067	A	G	A	A	G	T	T	A	A
12299	A	G	A	A	G	T	T	A	A
12299	A	G	A	A	G	T	T	A	A
12301	A	G	A	A	G	T	T	A	A
12300	A	G	A	A	G	T	T	A	A
C T T A T C C C A C A A T T A G G T T T T C T A T A T T C T A A A T T A A C T A A A A A A A T T G A Majority									
12360		12370		12380		12390		12400	
2344	C	T	T	A	T	C	C	C	A
2117	C	T	T	A	T	C	C	C	A
2349	C	T	T	A	T	C	C	C	A
2349	C	T	T	A	T	C	C	C	A
2351	C	T	T	A	T	C	C	C	A
2350	C	T	T	A	T	C	C	C	A
A C T T G A T A A T C G G T T G T C T C C G A C T G A A C A A A A G T T A T T G A T T A C C T T A T Majority									
12410		12420		12430		12440		12450	
2394	A	C	T	T	G	A	T	A	A
2167	A	C	T	T	G	A	T	A	A
2399	A	C	T	T	G	A	T	A	A
2399	A	C	T	T	G	A	T	A	A
2401	A	C	T	T	G	A	T	A	A
2400	A	C	T	T	G	A	T	A	A
T A T T A C A T A C T A A A G G T T T A A T C A T T G A T A T G T A A G A A G T A A G T C A G C T A Majority									
12460		12470		12480		12490		12500	
2444	T	A	T	T	A	C	A	T	A
2217	T	A	T	T	A	C	A	T	A
2449	T	A	T	T	A	C	A	T	A
2449	T	A	T	T	A	C	A	T	A
2451	T	A	T	T	A	C	A	T	A
2450	T	A	T	T	A	C	A	T	A
A C C G A T C T T T C T A T T C T A A A A C T T A T A T T G T T G C T T T A G A A A T T T T A A A G Majority									
12510		12520		12530		12540		12550	
2494	A	C	C	G	A	T	C	T	T
2267	A	C	C	G	A	T	C	T	T
2499	A	C	C	G	A	T	C	T	T
2499	A	C	C	G	A	T	C	T	T
2501	A	C	C	G	A	T	C	T	T
2500	A	C	C	G	A	T	C	T	T
A G C G T G G A T G G C T T C A T A A T A A A C A G A A A T C T T A C C A A T T T G C G A A G C C A Majority									
12560		12570		12580		12590		12600	
2544	A	G	C	G	T	G	G	A	T
2317	A	G	C	G	T	G	G	A	T
2549	A	G	C	G	T	G	G	A	T
2549	A	G	C	G	T	G	G	A	T
2551	A	G	C	G	T	G	G	A	T
2550	A	G	C	G	T	G	G	A	T

FIGURE 18 AI

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A A A A A T A T G A T A T T T G A A G A G T C C A A A G A T C T A A T A G A T A G T C C A G T T A G Majority					
	12610	12620	12630	12640	12650
2594	A A A A A T A T G A T A T T T G A A G A G T C C A A A G A T C T A A T A G A T A G T C C A G T T A G				2603_all.seq
2367	A A A A A T A T G A T A T T T G A A G A G T C C A A A G A T C T A A T A G A T A G T C C A G T T A G				18rs21_all.seq
2599	A A A A A T A T G A T A T T T G A A G A G T C C A A A G A T C T A A T A G A T A G T C C A G T T A G				cohl_all.seq
2599	A A A A A T A T G A T A T T T G A A G A G T C C A A A G A T C T A A T A G A T A G T C C A G T T A G				cjb111_all.seq
2601	A A A A A T A T G A T A T T T G A A G A G T C C A A A G A T C T A A T A G A T A G T C C A G T T A G				nem316_all.seq
2600	A A A A A T A T G A T A T T T G A A G A G T C C A A A G A T C T A A T A G A T A G T C C A G T T A G				a909_all.seq
A G A A G C G T T G A T T A T A A G T G A T A A G G A T T T T C A A A A A T T A A A A C A A G A G C Majority					
	12660	12670	12680	12690	12700
2644	A G A A G C G T T G A T T A T A A G T G A T A A G G A T T T T C A A A A A T T A A A A C A A G A G C				2603_all.seq
2417	A G A A G C G T T G A T T A T A A G T G A T A A G G A T T T T C A A A A A T T A A A A C A A G A G C				18rs21_all.seq
2649	A G A A G C G T T G A T T A T A A G T G A T A A G G A T T T T C A A A A A T T A A A A C A A G A G C				cohl_all.seq
2649	A G A A G C G T T G A T T A T A A G T G A T A A G G A T T T T C A A A A A T T A A A A C A A G A G C				cjb111_all.seq
2651	A G A A G C G T T G A T T A T A A G T G A T A A G G A T T T T C A A A A A T T A A A A C A A G A G C				nem316_all.seq
2650	A G A A G C G T T G A T T A T A A G T G A T A A G G A T T T T C A A A A A T T A A A A C A A G A G C				a909_all.seq
T A T T A T T T T A A C C G A C T T A T T T T A A A G A C T T A T C A T A T C T A G G C T T G C T T Majority					
	12710	12720	12730	12740	12750
2694	T A T T A T T T T A A C C G A C T T A T T T T A A A G A C T T A T C A T A T C T A G G C T T G C T T				2603_all.seq
2467	T A T T A T T T T A A C C G A C T T A T T T T A A A G A C T T A T C A T A T C T A G G C T T G C T T				18rs21_all.seq
2699	T A T T A T T T T A A C C G A C T T A T T T T A A A G A C T T A T C A T A T C T A G G C T T G C T T				cohl_all.seq
2699	T A T T A T T T T A A C C G A C T T A T T T T A A A G A C T T A T C A T A T C T A G G C T T G C T T				cjb111_all.seq
2701	T A T T A T T T T A A C C G A C T T A T T T T A A A G A C T T A T C A T A T C T A G G C T T G C T T				nem316_all.seq
2700	T A T T A T T T T A A C C G A C T T A T T T T A A A G A C T T A T C A T A T C T A G G C T T G C T T				a909_all.seq
G A T G A T T C G G A A A A A T A C G G A G A C T A T A C T A T T T T C A A G G A A A A G A T A C A A Majority					
	12760	12770	12780	12790	12800
2744	G A T G A T T C G G A A A A A T A C G G A G A C T A T A C T A T T T T C A A G G A A A A G A T A C A A				2603_all.seq
2517	G A T G A T T C G G A A A A A T A C G G A G A C T A T A C T A T T T T C A A G G A A A A G A T A C A A				18rs21_all.seq
2749	G A T G A T T C G G A A A A A T A C G G A G A C T A T A C T A T T T T C A A G G A A A A G A T A C A A				cohl_all.seq
2749	G A T G A T T C G G A A A A A T A C G G A G A C T A T A C T A T T T T C A A G G A A A A G A T A C A A				cjb111_all.seq
2751	G A T G A T T C G G A A A A A T A C G G A G A C T A T A C T A T T T T C A A G G A A A A G A T A C A A				nem316_all.seq
2750	G A T G A T T C G G A A A A A T A C G G A G A C T A T A C T A T T T T C A A G G A A A A G A T A C A A				a909_all.seq
A A G T T T C G A A T C A A G T C T T C A A C T A T A C A T C C T T C A A A G T C A T C G G C T A G Majority					
	12810	12820	12830	12840	12850
2794	A A G T T T C G A A T C A A G T C T T C A A C T A T A C A T C C T T C A A A G T C A T C G G C T A G				2603_all.seq
2567	A A G T T T C G A A T C A A G T C T T C A A C T A T A C A T C C T T C A A A G T C A T C G G C T A G				18rs21_all.seq
2799	A A G T T T C G A A T C A A G T C T T C A A C T A T A C A T C C T T C A A A G T C A T C G G C T A G				cohl_all.seq
2799	A A G T T T C G A A T C A A G T C T T C A A C T A T A C A T C C T T C A A A G T C A T C G G C T A G				cjb111_all.seq
2801	A A G T T T C G A A T C A A G T C T T C A A C T A T A C A T C C T T C A A A G T C A T C G G C T A G				nem316_all.seq
2800	A A G T T T C G A A T C A A G T C T T C A A C T A T A C A T C C T T C A A A G T C A T C G G C T A G				a909_all.seq
A G A T T T G G A A T T A T G A A C C A A T C C C T T T G A T T A C T A G A A A A A T A A A T A G C Majority					
	12860	12870	12880	12890	12900
2844	A G A T T T G G A A T T A T G A A C C A A T C C C T T T G A T T A C T A G A A A A A T A A A T A G C				2603_all.seq
2617	A G A T T T G G A A T T A T G A A C C A A T C C C T T T G A T T A C T A G A A A A A T A A A T A G C				18rs21_all.seq
2849	A G A T T T G G A A T T A T G A A C C A A T C C C T T T G A T T A C T A G A A A A A T A A A T A G C				cohl_all.seq
2849	A G A T T T G G A A T T A T G A A C C A A T C C C T T T G A T T A C T A G A A A A A T A A A T A G C				cjb111_all.seq
2851	A G A T T T G G A A T T A T G A A C C A A T C C C T T T G A T T A C T A G A A A A A T A A A T A G C				nem316_all.seq
2850	A G A T T T G G A A T T A T G A A C C A A T C C C T T T G A T T A C T A G A A A A A T A A A T A G C				a909_all.seq
T T G G A G A G G C T A A C G T G A C A C T G G T T G A T C C A A T C T C G C T T T A T T T A A C A Majority					
	12910	12920	12930	12940	12950
2894	T T G G A G A G G C T A A C G T G A C A C T G G T T G A T C C A A T C T C G C T T T A T T T A A C A				2603_all.seq
2667	T T G G A G A G G C T A A C G T G A C A C T G G T T G A T C C A A T C T C G C T T T A T T T A A C A				18rs21_all.seq
2899	T T G G A G A G G C T A A C G T G A C A C T G G T T G A T C C A A T C T C G C T T T A T T T A A C A				cohl_all.seq
2899	T T G G A G A G G C T A A C G T G A C A C T G G T T G A T C C A A T C T C G C T T T A T T T A A C A				cjb111_all.seq
2901	T T G G A G A G G C T A A C G T G A C A C T G G T T G A T C C A A T C T C G C T T T A T T T A A C A				nem316_all.seq
2900	T T G G A G A G G C T A A C G T G A C A C T G G T T G A T C C A A T C T C G C T T T A T T T A A C A				a909_all.seq

FIGURE 18 AJ



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Alignment Report of A1-1 alignment, using J. Hein method with Weighted residue weight table.  
Thursday, July 29, 2004 5:46 PM

C T A A G A A T G A T G A A G A C C C T C G T A T T G A A G A A G A A G T T G A G C A G C T A G A A Majority	
12960 12970 12980 12990 13000	
12944	C T A A G A A T G A T G A A G A C C C T C G T A T T G A A G A A G A A G T T G A G C A G C T A G A A 2603_all.seq
12717	C T A A G A A T G A T G A A G A C C C T C G T A T T G A A G A A G A A G T T G A G C A G C T A G A A 18rs21_all.seq
12949	C T A A G A A T G A T G A A G A C C C T C G T A T T G A A G A A G A A G T T G A G C A G C T A G A A coh1_all.seq
12949	C T A A G A A T G A T G A A G A C C C T C G T A T T G A A G A A G A A G T T G A G C A G C T A G A A cjb111_all.seq
12951	C T A A G A A T G A T G A A G A C C C T C G T A T T G A A G A A G A A G T T G A G C A G C T A G A A nem316_all.seq
12950	C T A A G A A T G A T G A A G A C C C T C G T A T T G A A G A A G A A G T T G A G C A G C T A G A A a909_all.seq

G A T A A G A T

Majority

12994	G A T A A G A T
12767	G A T A A G A T
12999	G A T A A G A T
12999	G A
13000	A
13000	G A T A A G A T

2603\_all.seq  
18rs21\_all.seq  
coh1\_all.seq  
cjb111\_all.seq  
nem316\_all.seq  
a909\_all.seq

Decoration 'Decoration #1': Shade (with solid black) residues that differ from the Consensus.

FIGURE 18 AK

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		Majority
GGCCTTGTTCCGATGTTGATCCGATAACTCCTGGCTCATTAAATAGCCTG		
1	GGCCTTGTTCCGATGTTGATCCGATAACTCCTGGCTCATTAAATAGCCTG	2603_al2.seq
1	GGCCTTGTTCCGATGTTGATCCGATAACTCCTGGCTCATTAAATAGCCTG	18rs21_al2.seq
1	GGCCTTGTTCCGATGTTGATCCGATAACTCCTGGCTCATTAAATAGCCTG	515_al2.seq
1	GGCCTTGTTCCGATGTTGATCCGATAACTCCTGGCTCATTAAATAGCCTG	cjb111_al2.seq
1	GGCCTTGTTCCGATGTTGATCCGATAACTCCTGGCTCATTAAATAGCCTG	h36b_al2.seq
TTCGTAACGCTCTTTTAATTATCTCTAACTTAGCATGGGTATTGGTAAAAAT		Majority
51	TTCGTAACGCTCTTTTAATTATCTCTAACTTAGCATGGGTATTGGTAAAAAT	2603_al2.seq
51	TTCGTAACGCTCTTTTAATTATCTCTAACTTAGCATGGGTATTGGTAAAAAT	18rs21_al2.seq
51	TTCGTAACGCTCTTTTAATTATCTCTAACTTAGCATGGGTATTGGTAAAAAT	515_al2.seq
51	TTCGTAACGCTCTTTTAATTATCTCTAACTTAGCATGGGTATTGGTAAAAAT	cjb111_al2.seq
51	TTCGTAACGCTCTTTTAATTATCTCTAACTTAGCATGGGTATTGGTAAAAAT	h36b_al2.seq
TTTGA AAAATAGACTAAGTATTTTATTAACCTCAGGCCACTTTCTATGCATG		Majority
101	TTTGA AAAATAGACTAAGTATTTTATTAACCTCAGGCCACTTTCTATGCATG	2603_al2.seq
101	TTTGA AAAATAGACTAAGTATTTTATTAACCTCAGGCCACTTTCTATGCATG	18rs21_al2.seq
101	TTTGA AAAATAGACTAAGTATTTTATTAACCTCAGGCCACTTTCTATGCATG	515_al2.seq
101	TTTGA AAAATAGACTAAGTATTTTATTAACCTCAGGCCACTTTCTATGCATG	cjb111_al2.seq
101	TTTGA AAAATAGACTAAGTATTTTATTAACCTCAGGCCACTTTCTATGCATG	h36b_al2.seq
AAATCAATTTCTTTTATAGAAATTGTTTACGGAATAGGAGCTTCTGGAGCAAC		Majority
151	AAATCAATTTCTTTTATAGAAATTGTTTACGGAATAGGAGCTTCTGGAGCAAC	2603_al2.seq
151	AAATCAATTTCTTTTATAGAAATTGTTTACGGAATAGGAGCTTCTGGAGCAAC	18rs21_al2.seq
151	AAATCAATTTCTTTTATAGAAATTGTTTACGGAATAGGAGCTTCTGGAGCAAC	515_al2.seq
151	AAATCAATTTCTTTTATAGAAATTGTTTACGGAATAGGAGCTTCTGGAGCAAC	cjb111_al2.seq
151	AAATCAATTTCTTTTATAGAAATTGTTTACGGAATAGGAGCTTCTGGAGCAAC	h36b_al2.seq
TATAGCATCCCTGAAACCAGAAACTGTGCAAAAAGTGCACCCCTCCTCTAG		Majority
201	TATAGCATCCCTGAAACCAGAAACTGTGCAAAAAGTGCACCCCTCCTCTAG	2603_al2.seq
201	TATAGCATCCCTGAAACCAGAAACTGTGCAAAAAGTGCACCCCTCCTCTAG	18rs21_al2.seq
201	TATAGCATCCCTGAAACCAGAAACTGTGCAAAAAGTGCACCCCTCCTCTAG	515_al2.seq
201	TATAGCATCCCTGAAACCAGAAACTGTGCAAAAAGTGCACCCCTCCTCTAG	cjb111_al2.seq
201	TATAGCATCCCTGAAACCAGAAACTGTGCAAAAAGTGCACCCCTCCTCTAG	h36b_al2.seq
CAACTGTTCCGTCTCTGTTAGGACAGTCAAAAACCAGCATCTATAGGTAAT		Majority
251	CAACTGTTCCGTCTCTGTTAGGACAGTCAAAAACCAGCATCTATAGGTAAT	2603_al2.seq
251	CAACTGTTCCGTCTCTGTTAGGACAGTCAAAAACCAGCATCTATAGGTAAT	18rs21_al2.seq
251	CAACTGTTCCGTCTCTGTTAGGACAGTCAAAAACCAGCATCTATAGGTAAT	515_al2.seq
251	CAACTGTTCCGTCTCTGTTAGGACAGTCAAAAACCAGCATCTATAGGTAAT	cjb111_al2.seq
251	CAACTGTTCCGTCTCTGTTAGGACAGTCAAAAACCAGCATCTATAGGTAAT	h36b_al2.seq
TTAAATATTTTTTTCTCCAAAGAGTTCTCGATAATAATCATTAAATCGCACG		Majority
301	TTAAATATTTTTTTCTCCAAAGAGTTCTCGATAATAATCATTAAATCGCACG	2603_al2.seq
301	TTAAATATTTTTTTCTCCAAAGAGTTCTCGATAATAATCATTAAATCGCACG	18rs21_al2.seq
301	TTAAATATTTTTTTCTCCAAAGAGTTCTCGATAATAATCATTAAATCGCACG	515_al2.seq
301	TTAAATATTTTTTTCTCCAAAGAGTTCTCGATAATAATCATTAAATCGCACG	cjb111_al2.seq
301	TTAAATATTTTTTTCTCCAAAGAGTTCTCGATAATAATCATTAAATCGCACG	h36b_al2.seq
ATAACGTTTTTTTCATAGGATAAATTGTATCACAATTTTAACTAAAATAAAC		Majority
351	ATAACGTTTTTTTCATAGGATAAATTGTATCACAATTTTAACTAAAATAAAC	2603_al2.seq
351	ATAACGTTTTTTTCATAGGATAAATTGTATCACAATTTTAACTAAAATAAAC	18rs21_al2.seq
351	ATAACGTTTTTTTCATAGGATAAATTGTATCACAATTTTAACTAAAATAAAC	515_al2.seq
351	ATAACGTTTTTTTCATAGGATAAATTGTATCACAATTTTAACTAAAATAAAC	cjb111_al2.seq
351	ATAACGTTTTTTTCATAGGATAAATTGTATCACAATTTTAACTAAAATAAAC	h36b_al2.seq
TCACTACTACAATAAAACTAAAAAAGATTGGAACGTCAGTTAGTCCCAAT		Majority
401	TCACTACTACAATAAAACTAAAAAAGATTGGAACGTCAGTTAGTCCCAAT	2603_al2.seq
401	TCACTACTACAATAAAACTAAAAAAGATTGGAACGTCAGTTAGTCCCAAT	18rs21_al2.seq
401	TCACTACTACAATAAAACTAAAAAAGATTGGAACGTCAGTTAGTCCCAAT	515_al2.seq
401	TCACTACTACAATAAAACTAAAAAAGATTGGAACGTCAGTTAGTCCCAAT	cjb111_al2.seq
401	TCACTACTACAATAAAACTAAAAAAGATTGGAACGTCAGTTAGTCCCAAT	h36b_al2.seq

Figure 19



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	C T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C Majority									
	460		470		480		490		500	
451	C T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C 2603_al2.seq									
451	C T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C 18rs21_al2.seq									
451	C T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C 515_al2.seq									
451	C T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C cjb111_al2.seq									
451	C T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C h36b_al2.seq									
	G C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C Majority									
	510		520		530		540		550	
501	G C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C 2603_al2.seq									
501	G C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C 18rs21_al2.seq									
501	G C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C 515_al2.seq									
501	G C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C cjb111_al2.seq									
501	G C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C h36b_al2.seq									
	T G T C G G A A A A T G A A C C C C T A G G T A A A T A C G A G A T A A C C C A A T T A A A A A A A Majority									
	560		570		580		590		600	
551	T G T C G G A A A A T G A A C C C C T A G G T A A A T A C G A G A T A A C C C A A T T A A A A A A A 2603_al2.seq									
551	T G T C G G A A A A T G A A C C C C T A G G T A A A T A C G A G A T A A C C C A A T T A A A A A A A 18rs21_al2.seq									
551	T G T C G G A A A A T G A A C C C C T A G G T A A A T A C G A G A T A A C C C A A T T A A A A A A A 515_al2.seq									
551	T G T C G G A A A A T G A A C C C C T A G G T A A A T A C G A G A T A A C C C A A T T A A A A A A A cjb111_al2.seq									
551	T G T C G G A A A A T G A A C C C C T A G G T A A A T A C G A G A T A A C C C A A T T A A A A A A A h36b_al2.seq									
	T G A G C A A A C C C A A A G T A C C T T G G C A C A A C A G T T T C C A T A T A C T C T T A G G C Majority									
	610		620		630		640		650	
601	T G A G C A A A C C C A A A G T A C C T T G G C A C A A C A G T T T C C A T A T A C T C T T A G G C 2603_al2.seq									
601	T G A G C A A A C C C A A A G T A C C T T G G C A C A A C A G T T T C C A T A T A C T C T T A G G C 18rs21_al2.seq									
601	T G A G C A A A C C C A A A A T A C C T T G G C A C A A C A G T T T C C A T A T A C T A T T A G G C 515_al2.seq									
601	T G A G C A A A C C C A A A A T A C C T T G G C A C A A C A G T T T C C A T A T A C T C T T A G G C cjb111_al2.seq									
601	T G A G C A A A C C C A A A G T A C C T T G G C A C A A C A G T T T C C A T A T A C T C T T A G G C h36b_al2.seq									
	A T A T A G T A C T G C A A T A A A A T A A T A T A C T C C C A A A T A T C A T A A A T G T T C C Majority									
	660		670		680		690		700	
651	A T A T A G T A C T G C A A T A A A A T A A T A T A C T C C C A A A T A T C A T A A A T G T T C C 2603_al2.seq									
651	A T A T A G T A C T G C A A T A A A A T A A T A T A C T C C C A A A T A T C A T A A A T G T T C C 18rs21_al2.seq									
651	A T A T A G T A C T G C A A T A A A A T A A T A T A C T C C C A A A T A T C A T A A A T G T T C C 515_al2.seq									
651	A T A T A G T A C T G C A A T A A A A T A A T A T A C T C C C A A A T A T C A T A A A T G T T C C cjb111_al2.seq									
651	A T A T A G T A C T G C A A T A A A A T A A T A T A C T C C C A A A T A T C A T A A A T G T T C C h36b_al2.seq									
	C A T C G A G T G C C C A C T G G G A A A C G A A T A G C C A C C T G C A A A T A C T A A A T G G G Majority									
	710		720		730		740		750	
701	C A T C G A G T G C C C A C T G G G A A A C G A A T A G C C A C C T G C A A A T A C T A A A T G G G 2603_al2.seq									
701	C A T C G A G T G C C C A C T G G G A A A C G A A T A G C C A C C T G C A A A T A C T A A A T G G G 18rs21_al2.seq									
701	C A T C G A G T G C C C A C T G G G A A A C G A A T A G C C A C C T G C A A A T A C T A A A T G G G 515_al2.seq									
701	C A T C G A G T G C C C A C T G G G A A A C G A A T A G C C A C C T G C A A A T A C T A A A T G G G cjb111_al2.seq									
701	C A T C G A G T G C C C A C T G G G A A A C G A A T A G C C A C C T G C A A A T A C T A A A T G G G h36b_al2.seq									
	T T A A A G T T G G T C T T A C T C T T T T G A A A A A T A A G T T T T A A A G A A A G T A T A C A T Majority									
	760		770		780		790		800	
751	T T A A A G T T G G T C T T A C T C T T T T G A A A A A T A A G T T T T A A A G A A A G T A T A C A T 2603_al2.seq									
751	T T A A A G T T G G T C T T A C T C T T T T G A A A A A T A A G T T T T A A A G A A A G T A T A C A T 18rs21_al2.seq									
751	T T A A A G T T G G T C T T A C T C T T T T G A A A A A T A A G T T T T A A A G A A A G T A T A C A T 515_al2.seq									
751	T T A A A G T T G G T C T T A C T C T T T T G A A A A A T A A G T T T T A A A G A A A G T A T A C A T cjb111_al2.seq									
751	T T A A A G T T G G T C T T A C T C T T T T G A A A A A T A A G T T T T A A A G A A A G T A T A C A T h36b_al2.seq									
	A T A C C A G A G A T A A T A G C A T T T A C T G C G A T A A A T C T A G C T T G A G G A T A C C A Majority									
	810		820		830		840		850	
801	A T A C C A G A G A T A A T A G C A T T T A C T G C G A T A A A T C T A G C T T G A G G A T A C C A 2603_al2.seq									
801	A T A C C A G A G A T A A T A G C A T T T A C T G C G A T A A A T C T A G C T T G A G G A T A C C A 18rs21_al2.seq									
801	A T A C C A G A G A T A A T A G C A T T T A C T G C G A T A A A T C T A G C T T G A G G A T A C C A 515_al2.seq									
801	A T A C C A G A G A T A A T A G C A T T T A C T G C G A T A A A T C T A G C T T G A G G A T A C C A cjb111_al2.seq									
801	A T A C C A G A G A T A A T A G C A T T T A C T G C G A T A A A T C T A G C T T G A G G A T A C C A h36b_al2.seq									
	C T T C T T A A G G T A A C A G A A A G T G A C G C T C A T A A T C G C A A T A G C T A T C T G G C Majority									
	860		870		880		890		900	
851	C T T C T T A A G G T A A C A G A A A G T G A C G C T C A T A A T C G C A A T A G C T A T C T G G C 2603_al2.seq									
851	C T T C T T A A G G T A A C A G A A A G T G A C G C T C A T A A T C G C A A T A G C T A T C T G G C 18rs21_al2.seq									
851	C T T C T T A A G G T A A C A G A A A G T G A C G C T C A T A A T C G C A A T A G C T A T C T G G C 515_al2.seq									
851	C T T C T T A A G G T A A C A G A A A G T G A C G C T C A T A A T C G C A A T A G C T A T C T G G C cjb111_al2.seq									
851	C T T C T T A A G G T A A C A G A A A G T G A C G C T C A T A A T C G C A A T A G C T A T C T G G C h36b_al2.seq									

FIGURE 19A

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	TTACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGA Majority									
	910	920	930	940	950					
901	TTACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGA 2603_al2.seq									
901	TTACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGA 18rs21_al2.seq									
901	TTACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGA 515_al2.seq									
901	TTACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGA cjb111_al2.seq									
901	TTACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGA h36b_al2.seq									
	TTTGGCAACTGTCCTCTAACACTTTCTTGAATAGTTTGGTCAAATGAAAT Majority									
	960	970	980	990	1000					
951	TTTGGCAACTGTCCTCTAACACTTTCTTGAATAGTTTGGTCAAATGAAAT 2603_al2.seq									
951	TTTGGCAACTGTCCTCTAACACTTTCTTGAATAGTTTGGTCAAATGAAAT 18rs21_al2.seq									
951	TTTGGCAACTGTCCTCTAACACTTTCTTGAATAGTTTGGTCAAATGAAAT 515_al2.seq									
951	TTTGGCAACTGTCCTCTAACACTTTCTTGAATAGTTTGGTCAAATGAAAT cjb111_al2.seq									
951	TTTGGCAACTGTCCTCTAACACTTTCTTGAATAGTTTGGTCAAATGAAAT h36b_al2.seq									
	TACAGTGTGCGGGCCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGA Majority									
	1010	1020	1030	1040	1050					
1001	TACAGTGTGCGGGCCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGA 2603_al2.seq									
1001	TACAGTGTGCGGGCCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGA 18rs21_al2.seq									
1001	TACAGTGTGCGGGCCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGA 515_al2.seq									
1001	TACAGTGTGCGGGCCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGA cjb111_al2.seq									
1001	TACAGTGTGCGGGCCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGA h36b_al2.seq									
	TAATAGCAATAAATGCTTGAATAAGTTTACTATTTTGAACGAGATAACATT Majority									
	1060	1070	1080	1090	1100					
1051	TAATAGCAATAAATGCTTGAATAAGTTTACTATTTTGAACGAGATAACATT 2603_al2.seq									
1051	TAATAGCAATAAATGCTTGAATAAGTTTACTATTTTGAACGAGATAACATT 18rs21_al2.seq									
1051	TAATAGCAATAAATGCTTGAATAAGTTTACTATTTTGAACGAGATAACATT 515_al2.seq									
1051	TAATAGCAATAAATGCTTGAATAAGTTTACTATTTTGAACGAGATAACATT cjb111_al2.seq									
1051	TAATAGCAATAAATGCTTGAATAAGTTTACTATTTTGAACGAGATAACATT h36b_al2.seq									
	AGTCTTTTTTATATCTTTCTAATATTGGCAAAACAAGCCACGTAAGTTAGAT Majority									
	1110	1120	1130	1140	1150					
1101	AGTCTTTTTTATATCTTTCTAATATTGGCAAAACAAGCCACGTAAGTTAGAT 2603_al2.seq									
1101	AGTCTTTTTTATATCTTTCTAATATTGGCAAAACAAGCCACGTAAGTTAGAT 18rs21_al2.seq									
1101	AGTCTTTTTTATATCTTTCTAATATTGGCAAAACAAGCCACGTAAGTTAGAT 515_al2.seq									
1101	AGTCTTTTTTATATCTTTCTAATATTGGCAAAACAAGCCACGTAAGTTAGAT cjb111_al2.seq									
1101	AGTCTTTTTTATATCTTTCTAATATTGGCAAAACAAGCCACGTAAGTTAGAT h36b_al2.seq									
	AGAAAAACAATCGAAATTAAATTTCCCTCAACGATATTAAATGGAATAACC Majority									
	1160	1170	1180	1190	1200					
1151	AGAAAAACAATCGAAATTAAATTTCCCTCAACGATATTAAATGGAATAACC 2603_al2.seq									
1151	AGAAAAACAATCGAAATTAAATTTCCCTCAACGATATTAAATGGAATAACC 18rs21_al2.seq									
1151	AGAAAAACAATCGAAATTAAATTTCCCTCAACGATATTAAATGGAATAACC 515_al2.seq									
1151	AGAAAAACAATCGAAATTAAATTTCCCTCAACGATATTAAATGGAATAACC cjb111_al2.seq									
1151	AGAAAAACAATCGAAATTAAATTTCCCTCAACGATATTAAATGGAATAACC h36b_al2.seq									
	ATTGTTAAAAGGTAATTGCCCTACACCAATAAATGTTCTGATATCAAAGTT Majority									
	1210	1220	1230	1240	1250					
1201	ATTGTTAAAAGGTAATTGCCCTACACCAATAAATGTTCTGATATCAAAGTT 2603_al2.seq									
1201	ATTGTTAAAAGGTAATTGCCCTACACCAATAAATGTTCTGATATCAAAGTT 18rs21_al2.seq									
1201	ATTGTTAAAAGGTAATTGCCCTACACCAATAAATGTTCTGATATCAAAGTT 515_al2.seq									
1201	ATTGTTAAAAGGTAATTGCCCTACACCAATAAATGTTCTGATATCAAAGTT cjb111_al2.seq									
1201	ATTGTTAAAAGGTAATTGCCCTACACCAATAAATGTTCTGATATCAAAGTT h36b_al2.seq									
	AGCAAATATAGCATACAAAGGAATCGCAAAGACATAGTTGAGAGCTACCA Majority									
	1260	1270	1280	1290	1300					
1251	AGCAAATATAGCATACAAAGGAATCGCAAAGACATAGTTGAGAGCTACCA 2603_al2.seq									
1251	AGCAAATATAGCATACAAAGGAATCGCAAAGACATAGTTGAGAGCTACCA 18rs21_al2.seq									
1251	AGCAAATATAGCATACAAAGGAATCGCAAAGACATAGTTGAGAGCTACCA 515_al2.seq									
1251	AGCAAATATAGCATACAAAGGAATCGCAAAGACATAGTTGAGAGCTACCA cjb111_al2.seq									
1251	AGCAAATATAGCATACAAAGGAATCGCAAAGACATAGTTGAGAGCTACCA h36b_al2.seq									
	TAGATACGGTCAAGCTAACTGTACCAAATAGACTAGCTTTAATAAAATCT Majority									
	1310	1320	1330	1340	1350					
1301	TAGATACGGTCAAGCTAACTGTACCAAATAGACTAGCTTTAATAAAATCT 2603_al2.seq									
1301	TAGATACGGTCAAGCTAACTGTACCAAATAGACTAGCTTTAATAAAATCT 18rs21_al2.seq									
1301	TAGATACGGTCAAGCTAACTGTACCAAATAGACTAGCTTTAATAAAATCT 515_al2.seq									
1301	TAGATACGGTCAAGCTAACTGTACCAAATAGACTAGCTTTAATAAAATCT cjb111_al2.seq									
1301	TAGATACGGTCAAGCTAACTGTACCAAATAGACTAGCTTTAATAAAATCT h36b_al2.seq									

FIGURE 19B



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TTTGCACCTCTCTCTATTTTCCAGAAAAATAGCGAAACTTGTCTAAAAATAA Majority									
1360 1370 1380 1390 1400									
1351	TTTGCACCTCTCTCTATTTTCCAGAAAAATAGCGAAACTTGTCTAAAAATAA	2603_al2.seq							
1351	TTTGCACCTCTCTCTATTTTCCAGAAAAATAGCGAAACTTGTCTAAAAATAA	18rs21_al2.seq							
1351	TTTGCACCTCTCTCTATTTTCCAGAAAAATAGCGAAACTTGTCTAAAAATAA	515_al2.seq							
1351	TTTGCACCTCTCTCTATTTTCCAGAAAAATAGCGAAACTTGTCTAAAAATAA	cjb111_al2.seq							
1351	TTTGCACCTCTCTCTATTTTCCAGAAAAATAGCGAAACTTGTCTAAAAATAA	h36b_al2.seq							
AGCTAGAGCAACCATATTCATCGGTAAACCAATAAACGTTTCTGGACCAC Majority									
1410 1420 1430 1440 1450									
1401	AGCTAGAGCAACCATATTCATCGGTAAACCAATAAACGTTTCTGGACCAC	2603_al2.seq							
1401	AGCTAGAGCAACCATATTCATCGGTAAACCAATAAACGTTTCTGGACCAC	18rs21_al2.seq							
1401	AGCTAGAGCAACCATATTCATCGGTAAACCAATAAACGTTTCTGGACCAC	515_al2.seq							
1401	AGCTAGAGCAACCATATTCATCGGTAAACCAATAAACGTTTCTGGACCAC	cjb111_al2.seq							
1401	AGCTAGAGCAACCATATTCATCGGTAAACCAATAAACGTTTCTGGACCAC	h36b_al2.seq							
GATTAGCAAGTATAAAGTGAATCTTAATAAGAGTACACCATAA Majority									
1460 1470 1480 1490 1500									
1451	GATTAGCAAGTATAAAGTGAATCTTAATAAGAGTACACCATAA	2603_al2.seq							
1451	GATTAGCAAGTATAAAGTGAATCTTAATAAGAGTACACCATAA	18rs21_al2.seq							
1451	GATTAGCAAGTATAAAGTGAATCTTAATAAGAGTACACCATAA	515_al2.seq							
1451	GATTAGCAAGTATAAAGTGAATCTTAATAAGAGTACACCATAA	cjb111_al2.seq							
1451	GATTAGCAAGTATAAAGTGAATCTTAATAAGAGTACACCATAA	h36b_al2.seq							
CTTGATTTTCAAAATCAAATAAAATAAAAGCAACTAACATCGGAAGGATTGA Majority									
1510 1520 1530 1540 1550									
1501	CTTGATTTTCAAAATCAAATAAAATAAAAGCAACTAACATCGGAAGGATTGA	2603_al2.seq							
1501	CTTGATTTTCAAAATCAAATAAAATAAAAGCAACTAACATCGGAAGGATTGA	18rs21_al2.seq							
1501	CTTGATTTTCAAAATCAAATAAAATAAAAGCAACTAACATCGGAAGGATTGA	515_al2.seq							
1501	CTTGATTTTCAAAATCAAATAAAATAAAAGCAACTAACATCGGAAGGATTGA	cjb111_al2.seq							
1501	CTTGATTTTCAAAATCAAATAAAATAAAAGCAACTAACATCGGAAGGATTGA	h36b_al2.seq							
AAAATCAACCTTTTAAATAATCTGCTCCTGGTATTAAATGGAAATGAAACCA Majority									
1560 1570 1580 1590 1600									
1551	AAAATCAACCTTTTAAATAATCTGCTCCTGGTATTAAATGGAAATGAAACCA	2603_al2.seq							
1551	AAAATCAACCTTTTAAATAATCTGCTCCTGGTATTAAATGGAAATGAAACCA	18rs21_al2.seq							
1551	AAAATCAACCTTTTAAATAATCTGCTCCTGGTATTAAATGGAAATGAAACCA	515_al2.seq							
1551	AAAATCAACCTTTTAAATAATCTGCTCCTGGTATTAAATGGAAATGAAACCA	cjb111_al2.seq							
1551	AAAATCAACCTTTTAAATAATCTGCTCCTGGTATTAAATGGAAATGAAACCA	h36b_al2.seq							
TCATCAATACAAAAGATAAGGCAGAAAGAAATGGCGATTGTACACCATTTTA Majority									
1610 1620 1630 1640 1650									
1601	TCATCAATACAAAAGATAAGGCAGAAAGAAATGGCGATTGTACACCATTTTA	2603_al2.seq							
1601	TCATCAATACAAAAGATAAGGCAGAAAGAAATGGCGATTGTACACCATTTTA	18rs21_al2.seq							
1601	TCATCAATACAAAAGATAAGGCAGAAAGAAATGGCGATTGTACACCATTTTA	515_al2.seq							
1601	TCATCAATACAAAAGATAAGGCAGAAAGAAATGGCGATTGTACACCATTTTA	cjb111_al2.seq							
1601	TCATCAATACAAAAGATAAGGCAGAAAGAAATGGCGATTGTACACCATTTTA	h36b_al2.seq							
CGTGTATTTGTCTATAAAAAAATTCCTCCAATTTAAATAAATTGAAAGAAAG Majority									
1660 1670 1680 1690 1700									
1651	CGTGTATTTGTCTATAAAAAAATTCCTCCAATTTAAATAAATTGAAAGAAAG	2603_al2.seq							
1651	CGTGTATTTGTCTATAAAAAAATTCCTCCAATTTAAATAAATTGAAAGAAAG	18rs21_al2.seq							
1651	CGTGTATTTGTCTATAAAAAAATTCCTCCAATTTAAATAAATTGAAAGAAAG	515_al2.seq							
1651	CGTGTATTTGTCTATAAAAAAATTCCTCCAATTTAAATAAATTGAAAGAAAG	cjb111_al2.seq							
1651	CGTGTATTTGTCTATAAAAAAATTCCTCCAATTTAAATAAATTGAAAGAAAG	h36b_al2.seq							
CTCCAAAGGTAAGCGTATGTACGCGAAAAAA - CCTTTGTCTTCTCCCATC Majority									
1710 1720 1730 1740 1750									
1701	CTCCAAAGGTAAGCGTATGTACGCGAAAAAA - CCTTTGTCTTCTCCCATC	2603_al2.seq							
1701	CTCCAAAGGTAAGCGTATGTACGCGAAAAAA - CCTTTGTCTTCTCCCATC	18rs21_al2.seq							
1701	CTCCAAAGGTAAGCGTATGTACGCGAAAAAA - CCTTTGTCTTCTCCCATC	515_al2.seq							
1701	CTCCAAAGGTAAGCGTATGTACGCGAAAAAA - CCTTTGTCTTCTCCCATC	cjb111_al2.seq							
1701	CTCCAAAGGTAAGCGTATGTACGCGAAAAAA - CCTTTGTCTTCTCCCATC	h36b_al2.seq							
CAGACTTTTACTGTCTGGTTGTGGAATCTCACACATCAGCTTTTCGCTCGCG Majority									
1760 1770 1780 1790 1800									
1750	CAGACTTTTACTGTCTGGTTGTGGAATCTCACACATCAGCTTTTCGCTCGCG	2603_al2.seq							
1750	CAGACTTTTACTGTCTGGTTGTGGAATCTCACACATCAGCTTTTCGCTCGCG	18rs21_al2.seq							
1750	CAGACTTTTACTGTCTGGTTGTGGAATCTCACACATCAGCTTTTCGCTCGCG	515_al2.seq							
1750	CAGACTTTTACTGTCTGGTTGTGGAATCTCACACATCAGCTTTTCGCTCGCG	cjb111_al2.seq							
1750	CAGACTTTTACTGTCTGGTTGTGGAATCTCACACATCAGCTTTTCGCTCGCG	h36b_al2.seq							

FIGURE 19C

FIGURE 19D



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A A A A A T A T A G T T A G C C A G C T A A T T A A C C T T C A A C C T C A A T T T C T C C C T T A T Majority									
2260		2270		2280		2290		2300	
2250	A A A A A T A T A C T T A C C A G G T A A T A A A C C T T C A A C C T C A A T T T C T C C C T T A T 2603_al2.seq								
2250	A A A A A T A T A C T T A C C A G G T A A T A A A C C T T C A A C C T C A A T T T C T C C C T T A T 18rs21_al2.seq								
2251	A A A A A T A T A C T T A C C A G G T A A T A A A C C T T C A A C C T C A A T T T C T C C C T T A T 515_al2.seq								
2250	A A A A A T A T A C T T A C C A G G T A A T A A A C C T T C A A C C T C A A T T T C T C C C T T A T cjb111_al2.seq								
2250	A A A A A T A T A C T T A C C A G G T A A T A A A C C T T C A A C C T C A A T T T C T C C C T T A T h36b_al2.seq								
C A T C A G T T A C T A A T G A A G T A A T C C C A T C T T G A T C G G T C G T A A A T C G T C C A Majority									
2310		2320		2330		2340		2350	
2300	C A T C A G T T A C T A A T G A A G T A A T C C C A T C T T G A T C G G T C G T A A A T C G T C C A 2603_al2.seq								
2300	C A T C A G T T A C T A A T G A A G T A A T C C C A T C T T G A T C G G T C G T A A A T C G T C C A 18rs21_al2.seq								
2301	C A T C A G T T A C T A A T G A A G T A A T C C C A T C T T G A T C G G T C G T A A A T C G T C C A 515_al2.seq								
2300	C A T C A G T T A C T A A T G A A G T A A T C C C A T C T T G A T C G G T C G T A A A T C G T C C A cjb111_al2.seq								
2300	C A T C A G T T A C T A A T G A A G T A A T C C C A T C T T G A T C G G T C G T A A A T C G T C C A h36b_al2.seq								
T T T T T A A A G C G A A C T G G C T G A T T C T G G T T A T C G T A T A A T A C A A A T A T T A C Majority									
2360		2370		2380		2390		2400	
2350	T T T T T A A A G C G A A C T G G C T G A T T C T G G T T A T C G T A T A A T A C A A A T A T T A C 2603_al2.seq								
2350	T T T T T A A A G C G A A C T G G C T G A T T C T G G T T A T C G T A T A A T A C A A A T A T T A C 18rs21_al2.seq								
2351	T T T T T A A A G C G A A C T G G C T G A T T C T G G T T A T C G T A T A A T A C A A A T A T T A C 515_al2.seq								
2350	T T T T T A A A G C G A A C T G G C T G A T T C T G G T T A T C G T A T A A T A C A A A T A T T A C cjb111_al2.seq								
2350	T T T T T A A A G C G A A C T G G C T G A T T C T G G T T A T C G T A T A A T A C A A A T A T T A C h36b_al2.seq								
T C C G G A T A G C C T T T T C T T T A T C T T T C C T T C T T T T G T A T A T T T A A T A A G T T Majority									
2410		2420		2430		2440		2450	
2400	T C C G G A T A G C C T T T T C T T T A T C T T T C C T T C T T T T G T A T A T T T A A T A A G T T 2603_al2.seq								
2400	T C C G G A T A G C C T T T T C T T T A T C T T T C C T T C T T T T G T A T A T T T A A T A A G T T 18rs21_al2.seq								
2401	T C C G G A T A G C C T T T T C T T T A T C T T T C C T T C T T T T G T A T A T T T A A T A A G T T 515_al2.seq								
2400	T C C G G A T A G C C T T T T C T T T A T C T T T C C T T C T T T T G T A T A T T T A A T A A G T T cjb111_al2.seq								
2400	T C C G G A T A G C C T T T T C T T T A T C T T T C C T T C T T T T G T A T A T T T A A T A A G T T h36b_al2.seq								
T T A A T C G G C C T G T T T C A A C T T T T C G C T T A G G A T T T A T C T G T A A T T G A T T T Majority									
2460		2470		2480		2490		2500	
2450	T T A A T C G G C C T G T T T C A A C T T T T C G C T T A G G A T T T A T C T G T A A T T G A T T T 2603_al2.seq								
2450	T T A A T C G G C C T G T T T C A A C T T T T C G C T T A G G A T T T A T C T G T A A T T G A T T T 18rs21_al2.seq								
2451	T T A A T C G G C C T G T T T C A A C T T T T C G C T T A G G A T T T A T C T G T A A T T G A T T T 515_al2.seq								
2450	T T A A T C G G C C T G T T T C A A C T T T T C G C T T A G G A T T T A T C T G T A A T T G A T T T cjb111_al2.seq								
2450	T T A A T C G G C C T G T T T C A A C T T T T C G C T T A G G A T T T A T C T G T A A T T G A T T T h36b_al2.seq								
G A T A A C T T A T C A T C T G G T A A T T C A A T A T A A A A G G T A C T A T T G T T G A A A C Majority									
2510		2520		2530		2540		2550	
2500	G A T A A C T T A T C A T C T G G T A A T T C A A T A T A A A A G G T A C T A T T G T T G A A A C 2603_al2.seq								
2500	G A T A A C T T A T C A T C T G G T A A T T C A A T A T A A A A G G T A C T A T T G T T G A A A C 18rs21_al2.seq								
2501	G A T A A C T T A T C A T C T G G T A A T T C A A T A T A A A A G G T A C T A T T G T T G A A A C 515_al2.seq								
2500	G A T A A C T T A T C A T C T G G T A A T T C A A T A T A A A A G G T A C T A T T G T T G A A A C cjb111_al2.seq								
2500	G A T A A C T T A T C A T C T G G T A A T T C A A T A T A A A A G G T A C T A T T G T T G A A A C h36b_al2.seq								
G C T T T G A T C A G C T T T A T A A G C A C G A C C A A A G T A C G A A C C A T T T G G G A G T G Majority									
2560		2570		2580		2590		2600	
2550	G C T T T G A T C A G C T T T A T A A G C A C G A C C A A A G T A C G A A C C A T T T G G G A G T G 2603_al2.seq								
2550	G C T T T G A T C A G C T T T A T A A G C A C G A C C A A A G T A C G A A C C A T T T G G G A G T G 18rs21_al2.seq								
2551	G C T T T G A T C A G C T T T A T A A G C A C G A C C A A A G T A C G A A C C A T T T G G G A G T G 515_al2.seq								
2550	G C T T T G A T C A G C T T T A T A A G C A C G A C C A A A G T A C G A A C C A T T T G G G A G T G cjb111_al2.seq								
2550	G C T T T G A T C A G C T T T A T A A G C A C G A C C A A A G T A C G A A C C A T T T G G G A G T G h36b_al2.seq								
C T A T C T T T G T C T G A C C A T T A G T A T C A G T A G G A G A A G T C A A G A T A C T C T T A Majority									
2610		2620		2630		2640		2650	
2600	C T A T C T T T G T C T G A C C A T T A G T A T C A G T A G G A G A A G T C A A G A T A C T C T T A 2603_al2.seq								
2600	C T A T C T T T G T C T G A C C A T T A G T A T C A G T A G G A G A A G T C A A G A T A C T C T T A 18rs21_al2.seq								
2601	C T A T C T T T G T C T G A C C A T T A G T A T C A G T A G G A G A A G T C A A G A T A C T C T T A 515_al2.seq								
2600	C T A T C T T T G T C T G A C C A T T A G T A T C A G T A G G A G A A G T C A A G A T A C T C T T A cjb111_al2.seq								
2600	C T A T C T T T G T C T G A C C A T T A G T A T C A G T A G G A G A A G T C A A G A T A C T C T T A h36b_al2.seq								
T A C T T C T G G T T C A A T T C G C T A T C T G T C A T T T G G C T C A A T A A A T C A A C T T T Majority									
2660		2670		2680		2690		2700	
2650	T A C T T C T G G T T C A A T T C G C T A T C T G T C A T T T G G C T C A A T A A A T C A A C T T T 2603_al2.seq								
2650	T A C T T C T G G T T C A A T T C G C T A T C T G T C A T T T G G C T C A A T A A A T C A A C T T T 18rs21_al2.seq								
2651	T A C T T C T G G T T C A A T T C G C T A T C T G T C A T T T G G C T C A A T A A A T C A A C T T T 515_al2.seq								
2650	T A C T T C T G G T T C A A T T C G C T A T C T G T C A T T T G G C T C A A T A A A T C A A C T T T cjb111_al2.seq								
2650	T A C T T C T G G T T C A A T T C G C T A T C T G T C A T T T G G C T C A A T A A A T C A A C T T T h36b_al2.seq								

FIGURE 19E

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TAAGTTGTCAGTCAACAGTCCATAAAACGATAAGAAATCCCCCTCCTCTGTAG Majority				
	2710	2720	2730	2740 2750
2700	TAAGTTGTCAGTCAACAGTCCATAAAACGATAAGAAATCCCCCTCCTCTGTAG 2603_a12.seq			
2700	TAAGTTGTCAGTCAACAGTCCATAAAACGATAAGAAATCCCCCTCCTCTGTAG 18rs21_a12.seq			
2701	TAAGTTGTCAGTCAACAGTCCATAAAACGATAAGAAATCCCCCTCCTCTGTAG 515_a12.seq			
2700	TAAGTTGTCAGTCAACAGTCCATAAAACGATAAGAAATCCCCCTCCTCTGTAG cjb111_a12.seq			
2700	TAAGTTGTCAGTCAACAGTCCATAAAACGATAAGAAATCCCCCTCCTCTGTAG h36b_a12.seq			
TATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCA Majority				
	2760	2770	2780	2790 2800
2750	TATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCA 2603_a12.seq			
2750	TATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCA 18rs21_a12.seq			
2751	TATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCA 515_a12.seq			
2750	TATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCA cjb111_a12.seq			
2750	TATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCA h36b_a12.seq			
GCATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAA Majority				
	2810	2820	2830	2840 2850
2800	GCATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAA 2603_a12.seq			
2800	GCATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAA 18rs21_a12.seq			
2801	GCATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAA 515_a12.seq			
2800	GCATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAA cjb111_a12.seq			
2800	GCATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAA h36b_a12.seq			
GCAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATT Majority				
	2860	2870	2880	2890 2900
2850	GCAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATT 2603_a12.seq			
2850	GCAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATT 18rs21_a12.seq			
2851	GCAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATT 515_a12.seq			
2850	GCAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATT cjb111_a12.seq			
2850	GCAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATT h36b_a12.seq			
TCCCTTTTCTCTTTTCTCTCTTTTAAATTTTTCGTTTTTAAATATAATAGTAAAG Majority				
	2910	2920	2930	2940 2950
2900	TCCCTTTTCTCTTTTCTCTCTTTTAAATTTTTCGTTTTTAAATATAATAGTAAAG 2603_a12.seq			
2900	TCCCTTTTCTCTTTTCTCTCTTTTAAATTTTTCGTTTTTAAATATAATAGTAAAG 18rs21_a12.seq			
2901	TCCCTTTTCTCTTTTCTCTCTTTTAAATTTTTCGTTTTTAAATATAATAGTAAAG 515_a12.seq			
2900	TCCCTTTTCTCTTTTCTCTCTTTTAAATTTTTCGTTTTTAAATATAATAGTAAAG cjb111_a12.seq			
2900	TCCCTTTTCTCTTTTCTCTCTTTTAAATTTTTCGTTTTTAAATATAATAGTAAAG h36b_a12.seq			
CGACTAATATAAGAATAAAGTAGGATTGATAAGAGGAAATAAAGTTTATAG Majority				
	2960	2970	2980	2990 3000
2950	CGACTAATATAAGAATAAAGTAGGATTGATAAGAGGAAATAAAGTTTATAG 2603_a12.seq			
2950	CGACTAATATAAGAATAAAGTAGGATTGATAAGAGGAAATAAAGTTTATAG 18rs21_a12.seq			
2951	CGACTAATATAAGAATAAAGTAGGATTGATAAGAGGAAATAAAGTTTATAG 515_a12.seq			
2950	CGACTAATATAAGAATAAAGTAGGATTGATAAGAGGAAATAAAGTTTATAG cjb111_a12.seq			
2950	CGACTAATATAAGAATAAAGTAGGATTGATAAGAGGAAATAAAGTTTATAG h36b_a12.seq			
TGTGTTTGGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTAC Majority				
	3010	3020	3030	3040 3050
3000	TGTGTTTGGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTAC 2603_a12.seq			
3000	TGTGTTTGGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTAC 18rs21_a12.seq			
3001	TGTGTTTGGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTAC 515_a12.seq			
3000	TGTGTTTGGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTAC cjb111_a12.seq			
3000	TGTGTTTGGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTAC h36b_a12.seq			
ATACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAAT Majority				
	3060	3070	3080	3090 3100
3050	ATACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAAT 2603_a12.seq			
3050	ATACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAAT 18rs21_a12.seq			
3051	ATACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAAT 515_a12.seq			
3050	ATACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAAT cjb111_a12.seq			
3050	ATACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAAT h36b_a12.seq			
AAGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAAT Majority				
	3110	3120	3130	3140 3150
3100	AAGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAAT 2603_a12.seq			
3100	AAGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAAT 18rs21_a12.seq			
3101	AAGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAAT 515_a12.seq			
3100	AAGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAAT cjb111_a12.seq			
3100	AAGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAAT h36b_a12.seq			

FIGURE 19F



A A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A										Majority
3160 3170 3180 3190 3200										
3150	A A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A									2603_al2.seq
3150	A A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A									18rs21_al2.seq
3151	A A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A									515_al2.seq
3150	A A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A									cjb111_al2.seq
3150	A A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A									h36b_al2.seq
G G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A										Majority
3210 3220 3230 3240 3250										
3200	G G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A									2603_al2.seq
3200	G G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A									18rs21_al2.seq
3201	G G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A									515_al2.seq
3200	G G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A									cjb111_al2.seq
3200	G G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A									h36b_al2.seq
G T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T										Majority
3260 3270 3280 3290 3300										
3250	G T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T									2603_al2.seq
3250	G T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T									18rs21_al2.seq
3251	G T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T									515_al2.seq
3250	G T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T									cjb111_al2.seq
3250	G T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T									h36b_al2.seq
G T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C										Majority
3310 3320 3330 3340 3350										
3300	G T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C									2603_al2.seq
3300	G T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C									18rs21_al2.seq
3301	G T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C									515_al2.seq
3300	G T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C									cjb111_al2.seq
3300	G T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C									h36b_al2.seq
A A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A										Majority
3360 3370 3380 3390 3400										
3350	A A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A									2603_al2.seq
3350	A A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A									18rs21_al2.seq
3351	A A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A									515_al2.seq
3350	A A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A									cjb111_al2.seq
3350	A A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A									h36b_al2.seq
T A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T T C A										Majority
3410 3420 3430 3440 3450										
3400	T A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T T C A									2603_al2.seq
3400	T A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T T C A									18rs21_al2.seq
3401	T A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T T C A									515_al2.seq
3400	T A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T T C A									cjb111_al2.seq
3400	T A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T T C A									h36b_al2.seq
G C A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T C T T T T T C										Majority
3460 3470 3480 3490 3500										
3450	G C A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C									2603_al2.seq
3450	G C A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C									18rs21_al2.seq
3451	G C A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C									515_al2.seq
3450	G C A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C									cjb111_al2.seq
3450	G C A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C									h36b_al2.seq
A G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T										Majority
3510 3520 3530 3540 3550										
3500	A G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T									2603_al2.seq
3500	A G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T									18rs21_al2.seq
3501	A G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T									515_al2.seq
3500	A G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T									cjb111_al2.seq
3500	A G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T									h36b_al2.seq
T T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T										Majority
3560 3570 3580 3590 3600										
3550	T T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T									2603_al2.seq
3550	T T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T									18rs21_al2.seq
3551	T T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T									515_al2.seq
3550	T T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T									cjb111_al2.seq
3550	T T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T									h36b_al2.seq

FIGURE 19G

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		T G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A										Majority
		3610		3620		3630		3640		3650		
3600	T G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A	2603_al2.seq										
3600	T G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A	18rs21_al2.seq										
3601	T G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A	515_al2.seq										
3600	T G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A	cjb111_al2.seq										
3600	T G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A	h36b_al2.seq										
		A T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A										Majority
		3660		3670		3680		3690		3700		
3650	A T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A	2603_al2.seq										
3650	A T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A	18rs21_al2.seq										
3651	A T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A A A A T A	515_al2.seq										
3650	A T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A	cjb111_al2.seq										
3650	A T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A	h36b_al2.seq										
		G T A T C A A G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G										Majority
		3710		3720		3730		3740		3750		
3700	G T A T C A A G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G	2603_al2.seq										
3700	G T A T C A A G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G	18rs21_al2.seq										
3701	A T A T C A A A C C T A A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G	515_al2.seq										
3700	G T A T C A A G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G	cjb111_al2.seq										
3700	G T A T C A A G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G	h36b_al2.seq										
		T T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T										Majority
		3760		3770		3780		3790		3800		
3750	T T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T	2603_al2.seq										
3750	T T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T	18rs21_al2.seq										
3751	T T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T	515_al2.seq										
3750	T T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T	cjb111_al2.seq										
3750	T T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T	h36b_al2.seq										
		G C T A A A C G A T T T T T T T G A C G T T G A C G T T T T A T T A A C C A A A G T A A C C A A G C										Majority
		3810		3820		3830		3840		3850		
3800	G C T A A A C G A T T T T T T T G A C G T T G A C G T T T T A T T A A C C A A A G T A A C C A A G C	2603_al2.seq										
3800	G C T A A A C G A T T T T T T T G A C G T T G A C G T T T T A T T A A C C A A A G T A A C C A A G C	18rs21_al2.seq										
3801	G A T A A A C G A T T T T T T T G A C G T T G A C G T T T T A T T A A C C A A A G T A A C C A A G C	515_al2.seq										
3800	G C T A A A C G A T T T T T T T G A C G T T G A C G T T T T A T T A A C C A A A G T A A C C A A G C	cjb111_al2.seq										
3800	G C T A A A C G A T T T T T T T G A C G T T G A C G T T T T A T T A A C C A A A G T A A C C A A G C	h36b_al2.seq										
		A A T A A T A A C T A A A G A T A T A T A G A A A A G A T A T C T A T A A A T C G T G T T T A A A T										Majority
		3860		3870		3880		3890		3900		
3850	A A T A A T A A C T A A A G A T A T A T A G A A A A G A T A T C T A T A A A T C G T G T T T A A A T	2603_al2.seq										
3850	A A T A A T A A C T A A A G A T A T A T A G A A A A G A T A T C T A T A A A T C G T G T T T A A A T	18rs21_al2.seq										
3851	A A T A A T A A C T A A A G A T A T A T A G A A A A G A T A T C T A T A A A T C G T G T T T A A A T	515_al2.seq										
3850	A A T A A T A A C T A A A G A T A T A T A G A A A A G A T A T C T A T A A A T C G T G T T T A A A T	cjb111_al2.seq										
3850	A A T A A T A A C T A A A G A T A T A T A G A A A A G A T A T C T A T A A A T C G T G T T T A A A T	h36b_al2.seq										
		G A C C G T C T T T T A T T A A T T T T T C A T C A A T A G G A C C T T T A T A A G G G A T A C G A										Majority
		3910		3920		3930		3940		3950		
3900	G A C C G T C T T T T A T T A A T T T T T C A T C A A T A G G A C C T T T A T A A G G G A T A C G A	2603_al2.seq										
3900	G A C C G T C T T T T A T T A A T T T T T C A T C A A T A G G A C C T T T A T A A G G G A T A C G A	18rs21_al2.seq										
3901	G A C C G T C T T T T A T T A A T T T T T C A T C A A T A G G A C C T T T A T A A G G G A T A C G A	515_al2.seq										
3900	G A C C T T C T T T T A T T A A T T T T T C A T C A A T A G G A C C T T T A T A A G G G A T A C G A	cjb111_al2.seq										
3900	G A C C T T C T T T T A T T A A T T T T T C A T C A A T A G G A C C T T T A T A A G G G A T A C G A	h36b_al2.seq										
		T G T C C C C T T A C T A A A A G T C T G T G T G T A T T G A C A T A A T C G G G G T G C A A G T										Majority
		3960		3970		3980		3990		4000		
3950	T G T C C C C T T A C T A A A A G T C T G T G T G T A T T G A C A T A A T C G G G G T G C A A G T	2603_al2.seq										
3950	T G T C C C C T T A C T A A A A G T C T G T G T G T A T T G A C A T A A T C G G G G T G C A A G T	18rs21_al2.seq										
3951	T G A C C C C T T A C T A A A A G T C T G T G T G T A T T G A C A T A A T C G G G G T G C A A G T	515_al2.seq										
3950	T G T C C C C T T A C T A A A A G T C T G T G T G T A T T G A C C A T A A T C G G G G T G C A A G T	cjb111_al2.seq										
3950	T G T C C C C T T A C T A A A A G T C T G T G T G T A T T G A C C A T A A T C G G G G T G C A A G T	h36b_al2.seq										
		C A A T A A G G T T G C A T A A T C A T G T C C A G G A A C A A C C A A C A A A T C T G A A A A G T										Majority
		4010		4020		4030		4040		4050		
4000	C A A T A A G G T T G C A T A A T C A T G T C C A G G A A C A A C C A A C A A A T C T G A A A A G T	2603_al2.seq										
4000	C A A T A A G G T T G C A T A A T C A T G T C C A G G A A C A A C C A A C A A A T C T G A A A A G T	18rs21_al2.seq										
4001	C A A T A A G G T T G C A T A A T C A T G T C C A G G A A C A A C C A A C A A A T C T G A A A A G T	515_al2.seq										
4000	A A A T A A G G T T G C A T A A T C A T G T C C A G G A A C A A C C A A C A A A T C T G A A A A G T	cjb111_al2.seq										
4000	C A A T A A G G T T G C A T A A T C A T G T C C A G G A A C A A C C A A C A A A T C T G A A A A G T	h36b_al2.seq										

FIGURE 19H



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TATCGGGGTGTAACGACTTTTATCTGATCTACTTGATATGCTATCGTTTCT Majority				
4060	4070	4080	4090	4100
4050	TATCGGGGTGTAACGACTTTTATCTGATCTACTTGATATGCTATCGTTTCT	2603_al2.seq		
4050	TATCGGGGTGTAACGACTTTTATCTGATCTACTTGATATGCTATCGTTTCT	18rs21_al2.seq		
4051	TATCGGGGTGTAACGACTTTTATCTGATCTACTTGATATGCTATCGTTTCT	515_al2.seq		
4050	TATCGGGGTGTAACGACTTTTATCTGATCTACTTGATATGCTATCGTTTCT	cjb111_al2.seq		
4050	TATCGGGGTGTAACGACTTTTATCTGATCTACTTGATATGCTATCGTTTCT	h36b_al2.seq		
TTTATGTTTTTGAATATAAAACTTTATCTCCTTTTTTTTAACTTTTTTAAGGTT Majority				
4110	4120	4130	4140	4150
4100	TTTATGTTTTTGAATATAAAACTTTATCTCCTTTTTTTTAACTTTTTTAAGGTT	2603_al2.seq		
4100	TTTATGTTTTTGAATATAAAACTTTATCTCCTTTTTTTTAACTTTTTTAAGGTT	18rs21_al2.seq		
4101	TTTATGTTTTTGAATATAAAACTTTATCTCCTTTTTTTTAACTTTTTTAAGGTT	515_al2.seq		
4100	TTTATGTTTTTGAATATAAAACTTTATCTCCTTTTTTTTAACTTTTTTAAGGTT	cjb111_al2.seq		
4100	TTTATGTTTTTGAATATAAAACTTTATCTCCTTTTTTTTAACTTTTTTAAGGTT	h36b_al2.seq		
AGAAAAGAGTTCTTTTATCTGGAATTCTGCAATGCGCTGTTATAACAGTAT Majority				
4160	4170	4180	4190	4200
4150	AGAAAAGAGTTCTTTTATCTGGAATTCTGCAATGCGCTGTTATAACAGTAT	2603_al2.seq		
4150	AGAAAAGAGTTCTTTTATCTGGAATTCTGCAATGCGCTGTTATAACAGTAT	18rs21_al2.seq		
4151	AGAAAAGAGTTCTTTTATCTGGAATTCTGCAATGCGCTGTTATAACAGTAT	515_al2.seq		
4150	AGAAAAGAGTTCTTTTATCTGGAATTCTGCAATGCGCTGTTATAACAGTAT	cjb111_al2.seq		
4150	AGAAAAGAGTTCTTTTATCTGGAATTCTGCAATGCGCTGTTATAACAGTAT	h36b_al2.seq		
GTGTAATAATTGCCCTCCAATTGGAAGAGAGGTACCTTCTAAATGCCCTGCT Majority				
4210	4220	4230	4240	4250
4200	GTGTAATAATTGCCCTCCAATTGGAAGAGAGGTACCTTCTAAATGCCCTGCT	2603_al2.seq		
4200	GTGTAATAATTGCCCTCCAATTGGAAGAGAGGTACCTTCTAAATGCCCTGCT	18rs21_al2.seq		
4201	GTGTAATAATTGCCCTCCAATTGGAAGAGAGGTACCTTCTAAATGCCCTGCT	515_al2.seq		
4200	GTGTAATAATTGCCCTCCAATTGGAAGAGAGGTACCTTCTAAATGCCCTGCT	cjb111_al2.seq		
4200	GTGTAATAATTGCCCTCCAATTGGAAGAGAGGTACCTTCTAAATGCCCTGCT	h36b_al2.seq		
CCTTTAGATAGAACTTCTTGACTTGAACTGCAAAATATAGGGAGTTTTTTG Majority				
4260	4270	4280	4290	4300
4250	CCTTTAGATAGAACTTCTTGACTTGAACTGCAAAATATAGGGAGTTTTTTG	2603_al2.seq		
4250	CCTTTAGATAGAACTTCTTGACTTGAACTGCAAAATATAGGGAGTTTTTTG	18rs21_al2.seq		
4251	CCTTTAGATAGAACTTCTTGACTTGAACTGCAAAATATAGGGAGTTTTTTG	515_al2.seq		
4250	CCTTTAGATAGAACTTCTTGACTTGAACTGCAAAATATAGGGAGTTTTTTG	cjb111_al2.seq		
4250	CCTTTAGATAGAACTTCTTGACTTGAACTGCAAAATATAGGGAGTTTTTTG	h36b_al2.seq		
ACCTATCTTAGGAACCTGAAATTGTGCGGATTTTTTTCACCTTACCTCTAACA Majority				
4310	4320	4330	4340	4350
4300	ACCTATCTTAGGAACCTGAAATTGTGCGGATTTTTTTCACCTTACCTCTAACA	2603_al2.seq		
4300	ACCTATCTTAGGAACCTGAAATTGTGCGGATTTTTTTCACCTTACCTCTAACA	18rs21_al2.seq		
4301	ACCTATCTTAGGAACCTGAAATTGTGCGGATTTTTTTCACCTTACCTCTAACA	515_al2.seq		
4300	ACCTATCTTAGGAACCTGAAATTGTGCGGATTTTTTTCACCTTACCTCTAACA	cjb111_al2.seq		
4300	ACCTATCTTAGGAACCTGAAATTGTGCGGATTTTTTTCACCTTACCTCTAACA	h36b_al2.seq		
TACGGGGCGTACTCTGCTACCCCTTTTGAATTTCGTTTTTTCTCATAAGGA Majority				
4360	4370	4380	4390	4400
4350	TACGGGGCGTACTCTGCTACCCCTTTTGAATTTCGTTTTTTCTCATAAGGA	2603_al2.seq		
4350	TACGGGGCGTACTCTGCTACCCCTTTTGAATTTCGTTTTTTCTCATAAGGA	18rs21_al2.seq		
4351	TACGGGGCGTACTCTGCTACCCCTTTTGAATTTCGTTTTTTCTCATAAGGA	515_al2.seq		
4350	TACGGGGCGTACTCTGCTACCCCTTTTGAATTTCGTTTTTTCTCATAAGGA	cjb111_al2.seq		
4350	TACGGGGCGTACTCTGCTACCCCTTTTGAATTTCGTTTTTTCTCATAAGGA	h36b_al2.seq		
TCTTCAAGATGGACATTATTTAAAGAATCATTATAAGCTTGTGCTAGAGC Majority				
4410	4420	4430	4440	4450
4400	TCTTCAAGATGGACATTATTTAAAGAATCATTATAAGCTTGTGCTAGAGC	2603_al2.seq		
4400	TCTTCAAGATGGACATTATTTAAAGAATCATTATAAGCTTGTGCTAGAGC	18rs21_al2.seq		
4401	TCTTCAAGATGGACATTATTTAAAGAATCATTATAAGCTTGTGCTAGAGC	515_al2.seq		
4400	TCTTCAAGATGGACATTATTTAAAGAATCATTATAAGCTTGTGCTAGAGC	cjb111_al2.seq		
4400	TCTTCAAGATGGACATTATTTAAAGAATCATTATAAGCTTGTGCTAGAGC	h36b_al2.seq		
CATACGTCGATTGATTTCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAA Majority				
4460	4470	4480	4490	4500
4450	CATACGTCGATTGATTTCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAA	2603_al2.seq		
4450	CATACGTCGATTGATTTCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAA	18rs21_al2.seq		
4451	CATACGTCGATTGATTTCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAA	515_al2.seq		
4450	CATACGTCGATTGATTTCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAA	cjb111_al2.seq		
4450	CATACGTCGATTGATTTCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAA	h36b_al2.seq		

FIGURE 19I

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AATCCTGTTGTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACC Majority				
4510	4520	4530	4540	4550
4500	AATCCTGTTGTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACC	2603_al2.seq		
4500	AATCCTGTTGTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACC	18rs21_al2.seq		
4501	AATCCTGTTGTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACC	515_al2.seq		
4500	AATCCTGTTGTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACC	cjb111_al2.seq		
4500	AATCCTGTTGTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACC	h36b_al2.seq		
ACTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGG Majority				
4560	4570	4580	4590	4600
4550	ACTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGG	2603_al2.seq		
4550	ACTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGG	18rs21_al2.seq		
4551	ACTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGG	515_al2.seq		
4550	ACTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGG	cjb111_al2.seq		
4550	ACTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGG	h36b_al2.seq		
AAGATTGACTTCTTCTTTTT - - - TGTTTTTTTTGATGATTTTTTTTACTCT Majority				
4610	4620	4630	4640	4650
4600	AAGATTGACTTCTTCTTTTT TTTT TGTTTTTTTTGATGATTTTTTTTACTCT	2603_al2.seq		
4600	AAGATTGACTTCTTCTTTTT TTTT TGTTTTTTTTGATGATTTTTTTTACTCT	18rs21_al2.seq		
4601	AAGATTGACTTCTTCTTTTT TTTT TGTTTTTTTTGATGATTTTTTTTACTCT	515_al2.seq		
4600	AAGATTGACTTCTTCTTTTT TTTT TGTTTTTTTTGATGATTTTTTTTACTCT	cjb111_al2.seq		
4600	AAGATTGACTTCTTCTTTTT TTTT TGTTTTTTTTGATGATTTTTTTTACTCT	h36b_al2.seq		
TCACGTCATCTCCTAAATAATGGCTCTTGCTTATGATCTAAGAGTACTTC Majority				
4660	4670	4680	4690	4700
4650	TCACGTCATCTCCTAAATAATGGCTCTTGCTTATGATCTAAGAGTACTTC	2603_al2.seq		
4650	TCACGTCATCTCCTAAATAATGGCTCTTGCTTATGATCTAAGAGTACTTC	18rs21_al2.seq		
4648	TCACGTCATCTCCTAAATAATGGCTCTTGCTTATGATCTAAGAGTACTTC	515_al2.seq		
4647	TCACGTCATCTCCTAAATAATGGCTCTTGCTTATGATCTAAGAGTACTTC	cjb111_al2.seq		
4647	TCACGTCATCTCCTAAATAATGGCTCTTGCTTATGATCTAAGAGTACTTC	h36b_al2.seq		
TACTGAAATACCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACAT Majority				
4710	4720	4730	4740	4750
4700	TACTGAAATACCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACAT	2603_al2.seq		
4700	TACTGAAATACCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACAT	18rs21_al2.seq		
4698	TACTGAAATACCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACAT	515_al2.seq		
4697	TACTGAAATACCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACAT	cjb111_al2.seq		
4697	TACTGAAATACCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACAT	h36b_al2.seq		
CATCAAAGACTAGCCTTAAGCTTCTCTGATTGACGTCCTTTTCATGATAA Majority				
4760	4770	4780	4790	4800
4750	CATCAAAGACTAGCCTTAAGCTTCTCTGATTGACGTCCTTTTCATGATAA	2603_al2.seq		
4750	CATCAAAGACTAGCCTTAAGCTTCTCTGATTGACGTCCTTTTCATGATAA	18rs21_al2.seq		
4748	CATCAAAGACTAGCCTTAAGCTTCTCTGATTGACGTCCTTTTCATGATAA	515_al2.seq		
4747	CATCAAAGACTAGCCTTAAGCTTCTCTGATTGACGTCCTTTTCATGATAA	cjb111_al2.seq		
4747	CATCAAAGACTAGCCTTAAGCTTCTCTGATTGACGTCCTTTTCATGATAA	h36b_al2.seq		
CTACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTA Majority				
4810	4820	4830	4840	4850
4800	CTACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTA	2603_al2.seq		
4800	CTACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTA	18rs21_al2.seq		
4798	CTACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTA	515_al2.seq		
4797	CTACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTA	cjb111_al2.seq		
4797	CTACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTA	h36b_al2.seq		
CCAATACCACCTGTTTGTGGGATTGTTACCTTTTTTATTCTACTCGTTG Majority				
4860	4870	4880	4890	4900
4850	CCAATACCACCTGTTTGTGGGATTGTTACCTTTTTTATTCTACTCGTTG	2603_al2.seq		
4850	CCAATACCACCTGTTTGTGGGATTGTTACCTTTTTTATTCTACTCGTTG	18rs21_al2.seq		
4848	CCAATACCACCTGTTTGTGGGATTGTTACCTTTTTTATTCTACTCGTTG	515_al2.seq		
4847	CCAATACCACCTGTTTGTGGGATTGTTACCTTTTTTATTCTACTCGTTG	cjb111_al2.seq		
4847	CCAATACCACCTGTTTGTGGGATTGTTACCTTTTTTATTCTACTCGTTG	h36b_al2.seq		
CGCATCTTTTTTTTTTGTGCTGCTAGCAGCGTAGTCAATGTTACCTG - - - AAC Majority				
4910	4920	4930	4940	4950
4900	CGCATCTTTTTTTTTTGTGCTGCTAGCAGCGTAGTCAATGTTACCTG - - - AAC	2603_al2.seq		
4900	CGCATCTTTTTTTTTTGTGCTGCTAGCAGCGTAGTCAATGTTACCTG - - - AAC	18rs21_al2.seq		
4898	CGCATCTTTTTTTTTTGTGCTGCTAGCAGCGTAGTCAATGTTACCTG - - - AAC	515_al2.seq		
4897	CGCATCTTTTTTTTTTGTGCTGCTAGCAGCGTAGTCAATGTTACCTG - - - AAC	cjb111_al2.seq		
4897	CGCATCTTTTTTTTTTGTGCTGCTAGCAGCGTAGTCAATGTTACCTG - - - AAC	h36b_al2.seq		

FIGURE 19J



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CTTTTCTGTAAGCTTTGATTAAGTTCAAACTTAACATCACCTGCCAAT Majority  
4960 4970 4980 4990 5000

4944 CTGTTATGTAATGACCTTTGATTAACTACAAACTTAAATATACCTGCCAAG 2603\_a12.seq  
4944 CTGTTATGTAATGACCTTTGATTAACTACAAACTTAAATATACCTGCCAAG 18rs21\_a12.seq  
4948 CTTTACTGTAATGAAGTAGGTAATAACATIAAAGCAACATCACCTGATAAT 515\_a12.seq  
4947 CTTTGCTATATGATGTTGGCAGTTTACTTCAAAGTTTACATCACCTGACAAT cjb111\_a12.seq  
4944 CTTTCTGCTTTTGTAGCTGTTCGAACCTTTHAAATGAAACATCACCTGCTAAT h36b\_a12.seq

TTAGCATAACCTGCTGGAGCTTGTGTTTCTTCAAGGTTGTAAAGTACCTTC Majority  
5010 5020 5030 5040 5050

4994 TTAGCAAATCCTGCTGGAGCAAGTGTTTCTTCAAGGTTGTAAAGTACCTGTC 2603\_a12.seq  
4994 TTAGCAAATCCTGCTGGAGCAAGTGTTTCTTCAAGGTTGTAAAGTACCTGTC 18rs21\_a12.seq  
4998 TTAGCATAACCAAGTTGGTGGCTTGTGTTTCTTCAAGGTTGTAAAGTACCTTC 515\_a12.seq  
4997 GTGGCATAACCTGCTGGTGGCTTGTGTTTCTTCAAGGTTGTAAAGTACCTTTI cjb111\_a12.seq  
4994 TTAGCGTAACCAAGCTGGAGCTTGTGTTTCTTCAAGTTTATAATCACCAAGT h36b\_a12.seq

TGCAAGACCTGTAACCTTCAAATTGACCTTGATCGTTTGAAGTCAATTTAA Majority  
5060 5070 5080 5090 5100

5044 TGCAAGACCTGTAACCTTCAAATTGACCTTGATCGTTTGAAGTGTAGGTAA 2603\_a12.seq  
5044 TGCAAGACCTGTAACCTTCAAATTGACCTTGATCGTTTGAAGTGTAGGTAA 18rs21\_a12.seq  
5048 AGTCAAGCCAGTAAATTTCAAATTGACCTTTATCGTTAGAAATCAATTTAA 515\_a12.seq  
5047 ATCCAAACCAGTAAATTTCAAATTGACCAACCGGCGTTAGABATCAATTTAA cjb111\_a12.seq  
5044 TGCAAGACCGCTAACCTTCAAATTGACCGTTAGCGTTTGAAGTCAATTTAA h36b\_a12.seq

CAACTTTAGCAGC-TTTTTATCTGTTACCCACTCATAAGCTGTACGAGCT Majority  
5110 5120 5130 5140 5150

5094 TGGCTCTAGC- - -CTTATCTGTTAICCACTCATAAGCTGTACGAGCT 2603\_a12.seq  
5094 TGGCTCTAGC- - -CTTATCTGTTAICCACTCATAAGCTGTACGAGCT 18rs21\_a12.seq  
5098 CAACATTTTLAGCATTTTATCTTCAACCCAATTCGTATGAGTAGTTAGCT 515\_a12.seq  
5097 CAACATTTTLAGCATTTTATCTTCAACCCAATTCATATGAGTAGTTAGCT cjb111\_a12.seq  
5094 CAACGTTAGCATCTTCTTTATTAGTTACCCACTCHTAAGCTGTACGAGCT h36b\_a12.seq

TCAATGAAGGCTGCATTGTAAAGCTTCTTTGTTTGTAGTTTGTAGTTGCTTT Majority  
5160 5170 5180 5190 5200

5138 TCAATGAAGGCTGCATTGTAAATCTGCTTCTTTAGTTTGTAGTTTGTAGTTT 2603\_a12.seq  
5138 TCAATGAAGGCTGCATTGTAAATCTGCTTCTTTAGTTTGTAGTTTGTAGTTT 18rs21\_a12.seq  
5148 TTAACAAGGCATCATTTGTAAAGCTTTTGTGTTTTCAGTAAAGGTCTTT 515\_a12.seq  
5147 TTAACAAGGCATCATTTGTAAAGCTTTTGTGTTTTCAGTAAAGGTCTTT cjb111\_a12.seq  
5144 ACGATGAAGGCTGCATTGTAAAGCTTCTTTGAGGAGTTTGTAGTTTCTGCTTT h36b\_a12.seq

TGCTGTTTTTACCTTTTTTGACCTTTTTTGTCTTCTGCGAGACAACCTTGTTAT Majority  
5210 5220 5230 5240 5250

5188 TGCAGTAATTCCTTTTTTACCTTTTTTGTCTCTGTTGCAGACAACCTTGTTAT 2603\_a12.seq  
5188 TGCAGTAATTCCTTTTTTACCTTTTTTGTCTCTGTTGCAGACAACCTTGTTAT 18rs21\_a12.seq  
5198 TGCTGATTTTACCATGTTTGACCTTTCTTGTTTTCTTTAGTCAAAATTTTGT 515\_a12.seq  
5197 TGCTGTTTTTACCTTTTGTGCGCTTTCTTGTTTTCTTTAGTCAAGTCTGTTAT cjb111\_a12.seq  
5194 AGCTGTTTTACCTTTTAGTACCTTTTTTGTCTCTGCTGAAAGTTTGTGTTGT h36b\_a12.seq

AAGCTGCGATAGCTTTCATCTAAAGCTATTTTAGTAGTAGCTAAAGCTGTT Majority  
5260 5270 5280 5290 5300

5238 AAGCAAGCGATAGCTTTCATCTAAAGCTATTTTCTTAGTAAAGCTT 2603\_a12.seq  
5238 AAGCAAGCGATAGCTTTCATCTAAAGCTATTTTCTTAGTAAAGCTT 18rs21\_a12.seq  
5248 AAGCTTTAAGAGCATCTCAAAGCTGATTTAGTTGAAICTAGAGCAAGCT 515\_a12.seq  
5247 AAGCTTTAAGAGCTTTCATCTAAAGCTAGTTTAGTAAAGTCTTCTAGAGCTGCT cjb111\_a12.seq  
5244 AAGCTGCGATTTTAAAGCTCAAGAGCAAGTTTTAGCAATTTCTGTAAGCTGTT h36b\_a12.seq

TTTTGAGCTGCTTTTGTCTTCTGCTTTAAGTGCAAGGTATTTACCTGC Majority  
5310 5320 5330 5340 5350

5285 TTTTGAAGCTTTTGTGATTGATCTGCTTTAAGAGCAAGGTATTTACCTGC 2603\_a12.seq  
5285 TTTTGAAGCTTTTGTGATTGATCTGCTTTAAGAGCAAGGTATTTACCTGC 18rs21\_a12.seq  
5298 TTTTCTGCACTGTTTGCACAAGCTGATTTAGTGCCAAAGTATTTGCTTCT 515\_a12.seq  
5297 TTTTCTGCACTGTTTGCACAAGCTGATTTAGTGCCAAAGTATTTGCTTCT cjb111\_a12.seq  
5294 GTTTTGAAGCTGATTTGTGCTGCAAGATGTTTCAATGCAAGGTATTTACCAAGC h36b\_a12.seq

TGAGTTTTTAAACAACGAATTGTGTCACCTGCCAAGCGTTTCAGTTTCAGGTT Majority  
5360 5370 5380 5390 5400

5332 TGAGTTTTTACAACAAGAAATTGTGTCACCAAGCCAAACGGTCACTTT 2603\_a12.seq  
5332 TGAGTTTTTACAACAAGAAATTGTGTCACCAAGCCAAACGGTCACTTT 18rs21\_a12.seq  
5348 TTTTCTTAAACAAGGAAGGTAGCTTCTGCCAAAGCGTTCTTTTCTCAATTT 515\_a12.seq  
5347 TTTTCTTAAACAAGGAAGGTAGCTTCTGCCAAAGCGTTCTTCTTCTCAATTT cjb111\_a12.seq  
5344 TGAAATTTTAAACAACGAATTGTGTCACCTTCAAGCGTTCTGTTTCTAGTCT h36b\_a12.seq

**FIGURE 19K**



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C A T T T G T T T T G A C A A A T T T T C T T A C C A T T G A G T C A C A A C T T T T G G T T C A G T T Majority									
5410		5420		5430		5440		5450	
5379	C	A	T	T	G	T	T	T	G
5379	C	A	T	T	G	T	T	T	G
5395	T	A	T	T	G	T	T	T	G
5394	G	A	T	T	G	T	T	T	G
5394	A	A	T	T	G	T	T	T	G
G G G T T G A T T G G T G T T G G G T T A T C T G A A T C T T T G G T A T T G G T G A T G G T T A C Majority									
5460		5470		5480		5490		5500	
5429	G	G	G	T	T	G	A	T	G
5429	G	G	G	T	T	G	A	T	G
5445	G	G	G	T	T	G	A	T	G
5444	G	G	T	T	G	A	T	G	T
5444	G	G	A	T	T	G	A	T	G
A T T A C C A T T T T C A A G A G T T A T - - - T G C A G T A C C G T A A C C A G T A A C A C G T T Majority									
5510		5520		5530		5540		5550	
5479	T	T	A	C	C	A	T	T	T
5479	T	T	A	C	C	A	T	T	T
5495	A	A	C	C	A	T	T	T	T
5494	A	A	C	C	A	T	T	T	T
5494	A	C	T	A	C	C	A	T	T
C T G A G A T C A T G T A T G T T T T A T T A T C A T C C A G A C C A G T G A A T T T A C C T G C G Majority									
5560		5570		5580		5590		5600	
5526	C	T	G	A	G	A	T	C	A
5526	C	T	G	A	G	A	T	C	A
5545	C	G	A	T	A	A	G	G	G
5544	C	G	A	T	A	A	G	G	G
5541	C	T	G	A	G	A	T	C	A
A A G T T A C C A G A T A C T G T A A A T T T G A T A C C A T T A C C A A T G T C G A T T G T A C C Majority									
5610		5620		5630		5640		5650	
5573	A	A	G	T	T	A	C	C	A
5573	A	A	G	T	T	A	C	C	A
5595	A	A	G	T	T	A	C	C	A
5594	A	A	G	T	T	A	C	C	A
5591	A	A	G	T	T	A	C	C	A
A T T - - - A G G T - - - G T T T T T G T C A A T G A T A C T G A A G C A A C A G C T G T A T C A T Majority									
5660		5670		5680		5690		5700	
5623	T	T	T	T	T	T	T	T	T
5623	T	T	T	T	T	T	T	T	T
5645	A	T	T	T	T	T	T	T	T
5644	A	T	T	T	T	T	T	T	T
5641	A	T	T	T	T	T	T	T	T
T A T C T T T A A A T G T G T A A A C A A C G T T T A C A T T A T C T G G T T C A C T A C C T T C T Majority									
5710		5720		5730		5740		5750	
5667	T	A	T	C	T	T	T	A	A
5667	T	A	T	C	T	T	T	A	A
5689	T	A	T	C	T	T	T	A	A
5691	T	A	T	C	T	T	T	A	A
5688	T	A	T	C	T	T	T	A	A
G C C C A A G T T T T A G T A A C T G T T A T T T C A C C C T T T G A T G G T G T A A C T G G T A G Majority									
5760		5770		5780		5790		5800	
5717	G	A	C	C	A	A	G	T	T
5717	G	A	C	C	A	A	G	T	T
5718	G	C	C	A	A	G	T	T	T
5717	G	C	C	A	A	G	T	T	T
5738	T	C	C	C	A	A	G	T	T
T T C T T T C A T T T T C T T T A C C T G G T T T G T T A C C A T A G T C C A A T T T G A T A T C A T Majority									
5810		5820		5830		5840		5850	
5767	T	T	C	A	G	T	C	A	A
5767	T	T	C	A	G	T	C	A	A
5768	T	T	C	A	G	T	C	A	A
5767	T	T	C	A	G	T	C	A	A
5788	T	T	C	A	G	T	C	A	A

FIGURE 19L



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		TGGTTTCTGCAATTTCACAAATAAGCTGCAACCGTTAACAGTAGCTCTATAA																				Majority																											
		5860					5870					5880					5890					5900																											
5817	T	G	G	A	T	T	C	T	G	G	A	T	T	A	T	C	A	A	I	A	A	T	T	G	C	T	T	G	A	C	C	A	T	T	A	A	2603_al2.seq												
5817	T	G	G	A	T	T	C	T	G	G	A	T	T	A	T	C	A	A	I	A	A	T	T	G	C	T	T	G	A	C	C	A	T	T	A	A	18rs21_al2.seq												
5818	T	G	G	T	T	T	C	T	A	G	A	A	C	T	T	C	A	A	C	A	C	A	G	A	A	C	C	G	T	T	A	A	A	G	T	A	G	C	T	G	A	G	T	A	A	515_al2.seq			
5817	T	G	G	T	T	T	C	T	G	G	A	T	T	T	C	A	A	C	A	C	A	G	T	A	G	C	C	G	T	T	A	C	C	G	T	A	G	C	T	G	A	G	T	A	A	cjb111_al2.seq			
5838	T	G	T	T	T	C	T	G	G	T	T	T	G	T	C	A	T	C	A	A	T	A	G	C	C	G	A	C	C	G	T	T	A	C	T	G	T	T	G	C	T	A	T	A	T	h36b_al2.seq			
		GTCAATGTGAATTCAACATCTTTTGTTTTTTGCTGCTTTTTTCCACTTTTTC																				Majority																											
		5910					5920					5930					5940					5950																											
5867	G	T	C	A	A	T	G	T	G	A	A	T	T	C	A	A	I	A	T	C	A	G	C	T	G	T	T	T	T	T	T	C	C	A	A	T	T	T	G	C	2603_al2.seq								
5867	G	T	C	A	A	T	G	T	G	A	A	T	T	C	A	A	I	A	T	C	A	G	C	T	G	T	T	T	T	T	T	C	C	A	A	T	T	T	G	C	18rs21_al2.seq								
5868	G	T	G	A	T	T	C	T	G	A	T	T	T	C	A	A	C	A	T	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	515_al2.seq					
5867	G	T	G	A	T	T	C	T	G	A	T	T	T	C	A	A	C	A	T	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	cjb111_al2.seq					
5888	G	T	C	A	A	T	G	T	G	A	A	T	T	C	A	A	C	A	G	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	h36b_al2.seq					
		CAAACCTTTAGCTGTGAATTTTAATGTGAAACCACGGTCAATCAGTTCCAA																				Majority																											
		5960					5970					5980					5990					6000																											
5917	C	A	A	I	C	C	A	T	C	A	G	C	T	G	T	G	A	A	T	T	T	A	A	T	G	T	G	A	A	A	C	C	A	C	G	G	C	A	T	C	A	A	T	G	C	T	A	A	2603_al2.seq
5917	C	A	A	T	C	C	A	T	C	A	G	C	T	G	T	G	A	A	T	T	T	A	A	T	G	T	G	A	A	A	C	C	A	C	G	G	C	A	T	C	A	A	T	G	C	T	A	A	18rs21_al2.seq
5918	A	A	G	A	C	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	515_al2.seq	
5917	A	A	G	A	C	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	cjb111_al2.seq	
5938	C	A	A	A	C	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	h36b_al2.seq	
		GTTTGTAGTCTGTATCCTTAACAAAAGTTCTG-----CCGATGCTTTCA																				Majority																											
		6010					6020					6030					6040					6050																											
5967	G	T	T	A	T	A	G	T	C	T	G	T	A	T	C	C	T	T	A	A	C	A	A	A	G	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	2603_al2.seq	
5967	G	T	T	A	T	A	G	T	C	T	G	T	A	T	C	C	T	T	A	A	C	A	A	A	G	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	18rs21_al2.seq	
5968	G	T	T	T	G	T	A	A	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	515_al2.seq	
5967	G	T	T	T	G	T	A	A	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	cjb111_al2.seq	
5988	C	A	T	T	G	T	A	A	T	C	T	G	T	A	C	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	h36b_al2.seq	
		AGGTTAACAGTGTTACCCATTGTCAAACCATTTTGACATGCTATCTGTCCA																				Majority																											
		6060					6070					6080					6090					6100																											
6017	A	G	G	T	T	A	A	C	A	G	T	G	T	T	A	C	C	C	A	T	T	G	T	C	A	A	A	C	C	A	T	T	T	G	A	C	A	T	T	T	T	T	T	T	T	T	T	2603_al2.seq	
6017	A	G	G	T	T	A	A	C	A	G	T	G	T	T	A	C	C	C	A	T	T	G	T	C	A	A	A	C	C	A	T	T	T	G	A	C	A	T	T	T	T	T	T	T	T	T	T	18rs21_al2.seq	
6012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	515_al2.seq	
6011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	cjb111_al2.seq	
6032	A	A	G	T	T	A	A	C	A	G	T	G	T	T	A	C	C	C	A	T	T	G	T	C	A	A	A	C	C	A	T	T	T	G	A	C	A	T	T	T	T	T	T	T	T	T	T	T	h36b_al2.seq
		AACCAAGTTTTTGTATTTAGAACCTTTGTGAATTTTGTTTTAACTTTCGT																				Majority																											
		6110					6120					6130					6140					6150																											
6067	A	A	C	C	A	A	G	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	2603_al2.seq	
6067	A	A	C	C	A	A	G	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	18rs21_al2.seq	
6059	G	A	T	T	A	A	I	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	515_al2.seq	
6058	A	A	C	C	A	A	G	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	cjb111_al2.seq	
6082	A	G	C	A	A	G	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	h36b_al2.seq	
		AAGGTATAACTTTACCGATTTCAGCAGTAGCAGTTGCTTTGTCCTTTTGA																				Majority																											
		6160					6170					6180					6190					6200																											
6117	A	A	G	G	A	A	C	A	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	2603_al2.seq	
6117	A	A	G	G	A	A	C	A	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	18rs21_al2.seq	
6109	A	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	515_al2.seq	
6108	A	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	I	cjb111_al2.seq	
6132	A	A	G	G	A	A	C	A	C	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	T	h36b_al2.seq	
		TAGTTAGCATAATCTGCGCCAGCTGTCAAAGTCTATTAACA-CTGTCAA																				Majority																											
		6210					6220					6230					6240					6250																											
6167	T	A	A	T	T	A	G	C																																									



**FIGURE 19N**



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G T T C C T G G G A A G T T A C T T T T G T C A A A A T A G C T T G T G T T T G G A C A A T C T T Majority									
6760		6770		6780		6790		6800	
6713	G T T C C T G G G A A G T T A C T T T T G T	A A G A T T T G A T T T G G T T T G C A A A A T C T T	2603_al2.seq						
6713	G T T C C T G G G A A G T T A C T T T T G T	A A G A T T T G A T T T G G T T T G C A A A A T C T T	18rs21_al2.seq						
6621	G T A C C A G C A G T T A A A A G T T C A A A T G A G T T G A G G C A I G A C A A T C T T	515_al2.seq							
6620	G T A C C T T C A G T T A A A G T T T C A A A T G A G C T T G T G G C A I G A C A A T C T T	cjb111_al2.seq							
6668	A T T C C T G G G A A A T T T G G C T T A A C A A A T T A C G G A T G T T T G A A C A A T T T T	h36b_al2.seq							
G T G C A A G G T C A C T G T A T T A G T - - - T G T - - - - T G C T T C A T C C G C A A A C G Majority									
6810		6820		6830		6840		6850	
6763	G T G C A A A G T C A C T G T A T T A G T - - - T G T - - - - T G C T T C A T C C G C A A A C G	2603_al2.seq							
6763	G T G C A A A G T C A C T G T A T T A G T - - - T G T - - - - T G C T T C A T C C G C A A A C G	18rs21_al2.seq							
6668	G T G C A A G G T C A C T G T T T A G T A G T T T T - - - - T G C T T C T C C G C A A A C A	515_al2.seq							
6667	G T G C A A G G T C A C A G T A T T A G T - - - T G T - - - - T G C G T C G T C C G C A A A C G	cjb111_al2.seq							
6718	G T G A A G G T T A C T G T T T A G T - - - G A T A T T T T C A T T C A G C A G C A A A F A	h36b_al2.seq							
C T G G T G C A A C T G A G A A C A A T G A C G T T A A A G T C A G T A A C A A T G C C G A G A A C Majority									
6860		6870		6880		6890		6900	
6804	C T G G T G C A A C T G A G A G C A A T G A C G T T A A A G T C A G T A A C A A T G C C G A G A A C	2603_al2.seq							
6804	C T G G T G C A A C T G A G A G C A A T G A C G T T A A A G T C A G T A A C A A T G C C G A G A A C	18rs21_al2.seq							
6712	C T G G T G C A A C T G A G A A C A A T G A C G T T A A A G T C A G T A A C A A T G C C G A G A A C	515_al2.seq							
6708	C T G G T G C A A C T G A G A A A A T G A C G T T A A A G T C A A T A G C A G T G C G A G A A C	cjb111_al2.seq							
6765	C T G G A G C T A C T G A A A C A A G A T G T T A A A A T T A A T A A C A A T G C C G A G A A C	h36b_al2.seq							
A T T G C A A A A T A T T T T G T T G A T T C T T T T C A T T T C T A T C T C C T T C T T A T T T T A Majority									
6910		6920		6930		6940		6950	
6854	A T T G C A A A A T A T T T T G T T G A T T C T T T T C A T T T C T A T C T C C T T C T T A T T T T A	2603_al2.seq							
6854	A T T G C A A A A T A T T T T G T T G A T T C T T T T C A T T T C T A T C T C C T T C T T A T T T T A	18rs21_al2.seq							
6762	A T T G C A A A A T A T T T T G T T G A T T T T T T C A T T T C T A T C T C C T T C T T A T T T T A	515_al2.seq							
6758	A T T G T A A G A T A T T T T G T T G A T T T T T T C A T T T C T A T C T C C T T C T T A T T T A	cjb111_al2.seq							
6815	A T T G C A A A A T A T T T T G T T G A T T C T T T T C A T T T C T A T C T C C T T C T T A T T T T A	h36b_al2.seq							
G T T A A T C A A C A T G A T T A A T A A T A T G C G G A T T T T A A T A - C - A C C G C A G C A C Majority									
6960		6970		6980		6990		7000	
6904	G T T A A T C A A C A T G A T T A A T A A T A T G C G G A T T T T A A T A - C - A C C G C A G C A C	2603_al2.seq							
6904	G T T A A T C A A C A T G A T T A A T A A T A T G C G G A T T T T A A T A - C - A C C G C A G C A C	18rs21_al2.seq							
6812	G T T A A T C A A C A T G A T T A A T A A T A T G C G G A T T T T A A T A - C - A C C G C A G C A C	515_al2.seq							
6808	G T T A A T C A A C A T A A A T A A T A A T A C G G A T T A A A T A - C - A C C G C A G C A C	cjb111_al2.seq							
6865	G T T A A T C A A C A T A A A T A A T A A T A C G G A T T A A A T A - C - A C C G C A G C A C	h36b_al2.seq							
C A C T C C C T T C A A G T C A T G G A A T T T T A G T T A A T T A A T T A A G A A T A C T A A A G Majority									
7010		7020		7030		7040		7050	
6952	C A C T C C C T T C A A G T C A T G G A A T T T T A G T T A A T T A A T T A A G A A T A C T A A A G	2603_al2.seq							
6952	C A C T C C C T T C A A G T C A T G G A A T T T T A G T T A A T T A A T T A A G A A T A C T A A A G	18rs21_al2.seq							
6860	C A C T C C C T T C A A G T C A T G G A A T T T T A G T T A A T T A A T T A A G A A T A C T A A A G	515_al2.seq							
6857	C A C T C C T T T C - - - - - G A A T T A T A T T A A T T A A T T A A G A A T A C T A A A A	cjb111_al2.seq							
6914	C A C T C C T T T C A A A A A T A T - - - - - A T T A A T T A A T A A T G C T A A A G	h36b_al2.seq							
C G C A T A A T T T T T A A T C T T T T T T G A T G G A C A T A T C A C T A G A T T T C T T A T A C Majority									
7060		7070		7080		7090		7100	
7002	C G C A T A A T T T T T A A T C T T T T T T G A T G G A C A T A T C A C T A G A T T T C T T A T A C	2603_al2.seq							
7002	C G C A T A A T T T T T A A T C T T T T T T G A T G G A C A T A T C A C T A G A T T T C T T A T A C	18rs21_al2.seq							
6910	C G C A T A A T T T T T A A T C T T T T T T G A T G G A C A T A T C A C T A G A T T T C T T A T A C	515_al2.seq							
6899	C A C A T A A T T T T T A A T C T T T T T T G A T G G A C A T A T C A C T A G A T T T C T T A T A	cjb111_al2.seq							
6952	C A C A T A T T T T T T A A T C T T T T T T G A T G G A C A T A T C A C T A G A T T T C T T A T A	h36b_al2.seq							
C T T T T C C A A A T A T A A A T T C C A C C T G C A A T A G A C A T C A T A G C T C C A C C T A T Majority									
7110		7120		7130		7140		7150	
7052	C T T T T C C A A A T A T A A A T T C C A C C T G C A A T A G A C A T C A T A G C T C C A C C T A T	2603_al2.seq							
7052	C T T T T C C A A A T A T A A A T T C C A C C T G C A A T A G A C A T C A T A G C T C C A C C T A T	18rs21_al2.seq							
6960	C T T T T C C A A A T A T A A A T T C C A C C T G C A A T A G A C A T C A T A G C T C C A C C T A T	515_al2.seq							
6949	C T T T T C C A A A T A T A A A T T C C A C C T G C A A T A G A C A T C A T A G A T C C A C C T A T	cjb111_al2.seq							
7002	C T T T T C C A A A T A T A A A T T C C A C C T G C A A T A G A C A T C A T A G C T C C A C C T A T	h36b_al2.seq							
T A A A A T G A A A G A T A G A A T T C C T T T C C C A C C T G T C A T A G G A A T A A T T C C T T Majority									
7160		7170		7180		7190		7200	
7102	T A A A A T G A A A G A T A G A A T T C C T T T C C C A C C T G T C A T A G G A A T A A T T C C T T	2603_al2.seq							
7102	T A A A A T G A A A G A T A G A A T T C C T T T C C C A C C T G T C A T A G G A A T A A T T C C T T	18rs21_al2.seq							
7010	T A A A A T G A A A G A T A G A A T T C C T T T C C C A C C T G T C T A G G A A T A A T T C C T T	515_al2.seq							
6999	T A A A A T G A A A G A T A G A A T T C C T T T C C C A C C T G T C A T G G A A T A A T T C C T T	cjb111_al2.seq							
7052	T A A A A T G A A A G A T A G A A T T C C T T T C C C A C C T G T C A T G G A A T A A T T C C T T	h36b_al2.seq							

FIGURE 190

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T T G G T G G A A T A T G C G T G T T G C T A A T T A A A T G C T T G T C A C C T T C C T C A T G A Majority									
7210		7220		7230		7240		7250	
7152	T T G G T G G A A T A T G C G T G T T G C T A A T T A A A T G C T T G T C A C C T T C C T C A T G A 2603_al2.seq								
7152	T T G G T G G A A T A T G C G T G T T G C T A A T T A A A T G C T T G T C A C C T T C C T C A T G A 18rs21_al2.seq								
7060	T T G C T G G A A T A T G C G T G T T G C T A A T T A A A T G C T T G T C A C C T T C C T C A T G A 515_al2.seq								
7049	T T G G T G G A A T A T G C G T G T T G C T A A T T A A A T G C T T G T C A C C T T C C T C A T G A cjb111_al2.seq								
7102	T T G G T G G A A T A T G C G T G T T G C T A A T T A A A T G C T T G T C A C C T T C C T C A T G A h36b_al2.seq								
T A T T C A G A A A T C T G T T T A T T A A C A G C T A T T A T A T T T T T T A T C G A T C C T T T Majority									
7260		7270		7280		7290		7300	
7202	T A T T C A G A A A T C T G T T T A T T A A C A G C T A T T A T A T T T T T T A T C G A T C C T T T 2603_al2.seq								
7202	T A T T C A G A A A T C T G T T T A T T A A C A G C T A T T A T A T T T T T T A T C G A T C C T T T 18rs21_al2.seq								
7110	T A T T C A G A A A T C T G T T T A T T A A C A G C T A T T A T A T T T T T T A T C G A T C C T T T 515_al2.seq								
7099	T A T T C A G A A A T C T G T T T A T T A A C A G C T A T T A T A T T T T T T A T C G A T C C T T T cjb111_al2.seq								
7152	T A T T C A G A A A T C T G T T T A T T A A C A G C T A T T A T A T T T T T T A T C G A T C C T T T h36b_al2.seq								
A A C C A C T T C A A A A G T T A A A A T T G G T T T A T T A G T A A T T T T T T T G A T A A T C C T Majority									
7310		7320		7330		7340		7350	
7252	A A C C A C T T C A A A A G T T A A A A T T G G T T T A T T A G T A A T T T T T T T G A T A A T C C T 2603_al2.seq								
7252	A A C C A C T T C A A A A G T T A A A A T T G G T T T A T T A G T A A T T T T T T T G A T A A T C C T 18rs21_al2.seq								
7160	A A C C A C T T C A A A A G T T A A A A T T G G T T T A T T A G T A A T T T T T T T G A T A A T C C T 515_al2.seq								
7149	A A C A A C T T C A A A A G T T A A A A T T G G T T T A T T A G T A A T T T T T T T G A T A A T C C T cjb111_al2.seq								
7202	A A C A A C T T C A A A A G T T A A A A T T G G T T T A T T A G T A A T T T T T T T G A T A A T C C T h36b_al2.seq								
C C G G C G A A A C T G C T T C T A T T A A C T G A T A T T T G C C A T C T T T C A A A T C T T T T G Majority									
7360		7370		7380		7390		7400	
7302	C C G G C G A A A C T G C T T C T A T T A A C T G A T A T T T G C C A T C T T T C A A A T C T T T G 2603_al2.seq								
7302	C C G G C G A A A C T G C T T C T A T T A A C T G A T A T T T G C C A T C T T T C A A A T C T T T G 18rs21_al2.seq								
7210	C C G G C G A A A C T G C T T C T A T T A A C T G A T A T T T G C C A T C T T T C A A A T C T T T G 515_al2.seq								
7199	T C G G C G A A A C T G C T T C T A T T A A C T G A T A T T T G C C A T C T T T C A A A T C T T T G cjb111_al2.seq								
7252	T C G G C G A A A C T G C T T C T A T T A A C T G A T A T T T G C C A T C T T T C A A A T C T T T G h36b_al2.seq								
T A A G A A A T T T T T G C C G T T T T C T C C C G T C A C T A C T T T T G A A T T A T T A T T T T T Majority									
7410		7420		7430		7440		7450	
7352	T A A G A A A T T T T T G C C G T T T T C T C C C G T C A C T A C T T T T G A A T T A T T A T T T T T 2603_al2.seq								
7352	T A A G A A A T T T T T G C C G T T T T C T C C C G T C A C T A C T T T T G A A T T A T T A T T T T T 18rs21_al2.seq								
7260	T A A G A A A T T T T T G C C G T T T T C T C C C G T C A C T A C T T T T G A A T T A T T A T T T T T 515_al2.seq								
7249	T A A G A A A T T T T T G C C G T T T T C T C C C G T C A C T A C T T T T G A A T T A T T A T T T T T cjb111_al2.seq								
7302	T A A G A A A T T T T T G C C G T T T T C T C C C G T C A C T A C T T T T G A A T T A T T A T T T T T h36b_al2.seq								
T A T T G G T A A A T A A A G T T T A T A A T C T T C A T T A A A T T C T T G A A G T T C A A A C G Majority									
7460		7470		7480		7490		7500	
7402	T A T T G G T A A A T A A A G T T T A T A A T C T T C A T T A A A T T C T T G A A G T T C A A A C G 2603_al2.seq								
7402	T A T T G G T A A A T A A A G T T T A T A A T C T T C A T T A A A T T C T T G A A G T T C A A A C G 18rs21_al2.seq								
7310	T A T T G G T A A A T A A A G T T T A T A A T C T T C A T T A A A T T C T T G A A G T T C A A A C G 515_al2.seq								
7299	T A T T G G T A A A T A A A G T T T A T A A T C T T C A T T A A A T T C T T G A A G T T C A A A C G cjb111_al2.seq								
7352	T A T T G G T A A A T A A A G T T T A T A A T C T T C A T T A A A T T C T T G A A G T T C A A A C G h36b_al2.seq								
T A G C T C C T T T G A G A A G C A A C T T A T T A T T A T C T T T A T C A A C T T T T A T A A A T Majority									
7510		7520		7530		7540		7550	
7452	T A G C T C C T T T G A G A A G C A A C T T A T T A T T A T C T T T A T C A A C T T T T A T A A A T 2603_al2.seq								
7452	T A G C T C C T T T G A G A A G C A A C T T A T T A T T A T C T T T A T C A A C T T T T A T A A A T 18rs21_al2.seq								
7360	T A G C T C C T T T G A G A A G C A A C T T A T T A T T A T C T T T A T C A A C T T T T A T A A A T 515_al2.seq								
7349	T A G C T C C T T T G A G A A G C A A C T T A T T A T T A T C T T T A T C A A C T T T T T A A A A T cjb111_al2.seq								
7402	T A G C T C C T T T G A G A A G C A A C T T A T T A T T A T C T T T A T C A A C T T T T T A A A A T h36b_al2.seq								
T C A A T T T C A C C T A A C T T C T T C T C G T T T T T A A T C G T T A T T G T A G G A T A T T C Majority									
7560		7570		7580		7590		7600	
7502	T C A A T T T C A C C T A A C T T C T T C T C G T T T T T A A T C G T T A T T G T A G G A T A T T C 2603_al2.seq								
7502	T C A A T T T C A C C T A A C T T C T T C T C G T T T T T A A T C G T T A T T G T A G G A T A T T C 18rs21_al2.seq								
7410	T C A A T T T C A C C T A A C T T C T T C T C G T T T T T A A T C G T T A T T G T A G G A T A T T C 515_al2.seq								
7399	T C A A T T T C A C C T A A C T T C T T C T C G T T T T T A A T C G T T A T T G T A G G A T A T T C cjb111_al2.seq								
7452	T C A A T T T C A C C T A A C T T C T T C T C G T T T T T A A T C G T T A T T G T A G G A T A T T C h36b_al2.seq								
T C T C A C A T C A C G A A T T T T A G G G A T T G G A A A A T C T C T A A G T G T A T T A G G A T Majority									
7610		7620		7630		7640		7650	
7552	T C T C A C A T C A C G A A T T T T A G G G A T T G G A A A A T C T C T A A G T G T A T T A G G A T 2603_al2.seq								
7552	T C T C A C A T C A C G A A T T T T A G G G A T T G G A A A A T C T C T A A G T G T A T T A G G A T 18rs21_al2.seq								
7460	T C T C A C A T C A C G A A T T T T A G G G A T T G G A A A A T C T C T A A G T G T A T T A G G A T 515_al2.seq								
7449	T C T C A C A T C A C G A A T T T T A G G G A T T G G A A A A T C T C T A A G T G T A T T A G G A T cjb111_al2.seq								
7502	T C T C A C A T C A C G A A T T T T A G G G A T T G G A A A A T C T C T A A G T G T A T T A G G A T h36b_al2.seq								

FIGURE 19P



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CCTCTGACTTAGGATTCAATGTTGTTCTACCATTAGTGTTCATAGAATTTC Majority									
7660		7670		7680		7690		7700	
7602	CCTCTGACTTAGGATTCAATGTTGTTCTACCATTAGTGTTCATAGAATTTC	2603_al2.seq							
7602	CCTCTGACTTAGGATTCAATGTTGTTCTACCATTAGTGTTCATAGAATTTC	18rs21_al2.seq							
7510	CCTCTGACTTAGGATTCAATGTTGTTCTACCATTAGTGTTCATAGAATTTC	515_al2.seq							
7499	CCTCTGACTTAGGATTCAATGTTGTTCTACCATTAGTGTTCATAGAATTTC	cjb111_al2.seq							
7552	CCTCTGACTTAGGATTCAATGTTGTTCTACCATTAGTGTTCATAGAATTTC	h36b_al2.seq							
TTACTTTATAAACTGTCATCTAGTTTTCACATCATATGTCAGTGTACTTTT Majority									
7710		7720		7730		7740		7750	
7652	TTACTTTATAAACTGTCATCTAGTTTTCACATCATATGTCAGTGTACTTTT	2603_al2.seq							
7652	TTACTTTATAAACTGTCATCTAGTTTTCACATCATATGTCAGTGTACTTTT	18rs21_al2.seq							
7560	TTACTTTATAAACTGTCATCTAGTTTTCACATCATATGTCAGTGTACTTTT	515_al2.seq							
7549	TTACTTTATAAACTGTCATCTAGTTTTCACATCATATGTCAGTGTACTTTT	cjb111_al2.seq							
7602	TTACTTTATAAACTGTCATCTAGTTTTCACATCATATGTCAGTGTACTTTT	h36b_al2.seq							
TTGACCTTCTCCTAAATTCAAACTCTAACATAGAGTTTATTTCCGATGT Majority									
7760		7770		7780		7790		7800	
7702	TTGACCTTCTCCTAAATTCAAACTCTAACATAGAGTTTATTTCCGATGT	2603_al2.seq							
7702	TTGACCTTCTCCTAAATTCAAACTCTAACATAGAGTTTATTTCCGATGT	18rs21_al2.seq							
7610	TTGACCTTCTCCTAAATTCAAACTCTAACATAGAGTTTATTTCCGATGT	515_al2.seq							
7599	TTGACCTTCTCCTAAATTCAAACTCTAACATAGAGTTTATTTCCGATGT	cjb111_al2.seq							
7652	TTGACCTTCTCCTAAATTCAAACTCTAACATAGAGTTTATTTCCGATGT	h36b_al2.seq							
ATTCTAATTTAAACCCCTTAAGTATTCCACCATCATTATTAGGCCCAACA Majority									
7810		7820		7830		7840		7850	
7752	ATTCTAATTTAAACCCCTTAAGTATTCCACCATCATTATTAGGCCCAACA	2603_al2.seq							
7752	ATTCTAATTTAAACCCCTTAAGTATTCCACCATCATTATTAGGCCCAACA	18rs21_al2.seq							
7660	ATTCTAATTTAAACCCCTTAAGTATTCCACCATCATTATTAGGCCCAACA	515_al2.seq							
7649	ATTCTAATTTAAACCCCTTAAGTATTCCACCATCATTATTAGGCCCAACA	cjb111_al2.seq							
7702	ATTCTAATTTAAACCCCTTAAGTATTCCACCATCATTATTAGGCCCAACA	h36b_al2.seq							
GTTGCAATACCATCCTTCATTACACTTCCATCATTTCCTGTAAAGTATA Majority									
7860		7870		7880		7890		7900	
7802	GTTGCAATACCATCCTTCATTACACTTCCATCATTTCCTGTAAAGTATA	2603_al2.seq							
7802	GTTGCAATACCATCCTTCATTACACTTCCATCATTTCCTGTAAAGTATA	18rs21_al2.seq							
7710	GTTGCAATACCATCCTTCATTACACTTCCATCATTTCCTGTAAAGTATA	515_al2.seq							
7699	GTTGCAATACCATCCTTCATTACACTTCCATCATTTCCTGTAAAGTATA	cjb111_al2.seq							
7752	GTTGCAATACCATCCTTCATTACACTTCCATCATTTCCTGTAAAGTATA	h36b_al2.seq							
ATCACTTGGCTGTAAATGTTTGTCCATTACCAAGCTGTAAATTGATTTTAT Majority									
7910		7920		7930		7940		7950	
7852	ATCACTTGGCTGTAAATGTTTGTCCATTACCAAGCTGTAAATTGATTTTAT	2603_al2.seq							
7852	ATCACTTGGCTGTAAATGTTTGTCCATTACCAAGCTGTAAATTGATTTTAT	18rs21_al2.seq							
7760	ATCACTTGGCTGTAAATGTTTGTCCATTACCAAGCTGTAAATTGATTTTAT	515_al2.seq							
7749	ATCACTTGGCTGTAAATGTTTGTCCATTACCAAGCTGTAAATTGATTTTAT	cjb111_al2.seq							
7802	ATCACTTGGCTGTAAATGTTTGTCCATTACCAAGCTGTAAATTGATTTTAT	h36b_al2.seq							
CACCCATAGGATCTTCGATAGTTCCATTAAACAATTGAGTTTCTTTTGT Majority									
7960		7970		7980		7990		8000	
7902	CACCCATAGGATCTTCGATAGTTCCATTAAACAATTGAGTTTCTTTTGT	2603_al2.seq							
7902	CACCCATAGGATCTTCGATAGTTCCATTAAACAATTGAGTTTCTTTTGT	18rs21_al2.seq							
7810	CACCCATAGGATCTTCGATAGTTCCATTAAACAATTGAGTTTCTTTTGT	515_al2.seq							
7799	CACCCATAGGATCTTCGATAGTTCCATTAAACAATTGAGTTTCTTTTGT	cjb111_al2.seq							
7852	CACCCATAGGATCTTCGATAGTTCCATTAAACAATTGAGTTTCTTTTGT	h36b_al2.seq							
AAAATCGTTTCAAATTGTTGCTGAATTTTAGATAAAAATTTTCATTGTTAGA Majority									
8010		8020		8030		8040		8050	
7952	AAAATCGTTTCAAATTGTTGCTGAATTTTAGATAAAAATTTTCATTGTTAGA	2603_al2.seq							
7952	AAAATCGTTTCAAATTGTTGCTGAATTTTAGATAAAAATTTTCATTGTTAGA	18rs21_al2.seq							
7860	AAAATCGTTTCAAATTGTTGCTGAATTTTAGATAAAAATTTTCATTGTTAGA	515_al2.seq							
7849	AAAATCGTTTCAAATTGTTGCTGAATTTTAGATAAAAATTTTCATTGTTAGA	cjb111_al2.seq							
7902	AAAATCGTTTCAAATTGTTGCTGAATTTTAGATAAAAATTTTCATTGTTAGA	h36b_al2.seq							
TGTATCGGCTGAAGTTACGATAGGGGTGTAGTACTCAGGTTTGGGAAGAGA Majority									
8060		8070		8080		8090		8100	
8002	TGTATCGGCTGAAGTTACGATAGGGGTGTAGTACTCAGGTTTGGGAAGAGA	2603_al2.seq							
8002	TGTATCGGCTGAAGTTACGATAGGGGTGTAGTACTCAGGTTTGGGAAGAGA	18rs21_al2.seq							
7910	TGTATCGGCTGAAGTTACGATAGGGGTGTAGTACTCAGGTTTGGGAAGAGA	515_al2.seq							
7899	TGTATCGGCTGAAGTTACGATAGGGGTGTAGTACTCAGGTTTGGGAAGAGA	cjb111_al2.seq							
7952	TGTATCGGCTGAAGTTACGATAGGGGTGTAGTACTCAGGTTTGGGAAGAGA	h36b_al2.seq							

FIGURE 19Q

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		ACGACCTCATTAGTTCTGTGATTTCCTCCATCTGAAAGTTTAAAAGCTTCC Majority									
		8110		8120		8130		8140		8150	
8052		ACGACCTCATTAGTTCTGTGATTTCCTCCATCTGAAAGTTTAAAAGCTTCC 2603_a12.seq									
8052		ACGACCTCATTAGTTCTGTGATTTCCTCCATCTGAAAGTTTAAAAGCTTCC 18rs21_a12.seq									
7960		ACGACCTCATTAGTTCTGTGATTTCCTCCATCTGAAAGTTTAAAAGCTTCC 515_a12.seq									
7949		ACGACCTCATTAGTTCTGTGATTTCCTCCATCTGAAAGTTTAAAAGCTTCC cjb111_a12.seq									
8002		ACGACCTCATTAGTTCTGTGATTTCCTCCATCTGAAAGTTTAAAAGCTTCC h36b_a12.seq									
		TCTTTCAATTTTGTAAAAGTACCATCTTGATTTTCTTATACTCCTCATTT Majority									
		8160		8170		8180		8190		8200	
8102		TCTTTCAATTTTGTAAAAGTACCATCTTGATTTTCTTATACTCCTCATTT 2603_a12.seq									
8102		TCTTTCAATTTTGTAAAAGTACCATCTTGATTTTCTTATACTCCTCATTT 18rs21_a12.seq									
8010		TCTTTCAATTTTGTAAAAGTACCATCTTGATTTTCTTATACTCCTCATTT 515_a12.seq									
7999		TCTTTCAATTTTGTAAAAGTACCATCTTGATTTTCTTATACTCCTCATTT cjb111_a12.seq									
8052		TCTTTCAATTTTGTAAAAGTACCATCTTGATTTTCTTATACTCCTCATTT h36b_a12.seq									
		ATAAACTTGTCTAAAACCAGATATATCGATACCAAATTAATAATGTCAT Majority									
		8210		8220		8230		8240		8250	
8152		ATAAACTTGTCTAAAACCAGATATATCGATACCAAATTAATAATGTCAT 2603_a12.seq									
8152		ATAAACTTGTCTAAAACCAGATATATCGATACCAAATTAATAATGTCAT 18rs21_a12.seq									
8060		ATAAACTTGTCTAAAACCAGATATATCGATACCAAATTAATAATGTCAT 515_a12.seq									
8049		ATAAACTTGTCTAAAACCAGATATATCGATACCAAATTAATAATGTCAT cjb111_a12.seq									
8102		ATAAACTTGTCTAAAACCAGATATATCGATACCAAATTAATAATGTCAT h36b_a12.seq									
		AATTTTCTGTTTTAAACTATTTATATAAAGTTTGCTTGCTGTTCCATGT Majority									
		8260		8270		8280		8290		8300	
8202		AATTTTCTGTTTTAAACTATTTATATAAAGTTTGCTTGCTGTTCCATGT 2603_a12.seq									
8202		AATTTTCTGTTTTAAACTATTTATATAAAGTTTGCTTGCTGTTCCATGT 18rs21_a12.seq									
8110		AATTTTCTGTTTTAAACTATTTATATAAAGTTTGCTTGCTGTTCCATGT 515_a12.seq									
8099		AATTTTCTGTTTTAAACTATTTATATAAAGTTTGCTTGCTGTTCCATGT cjb111_a12.seq									
8152		AATTTTCTGTTTTAAACTATTTATATAAAGTTTGCTTGCTGTTCCATGT h36b_a12.seq									
		TCTTTCACTGCTCCATTTTCGATAAAATTGTACCTTTAGGGTAATTAAGATT Majority									
		8310		8320		8330		8340		8350	
8252		TCTTTCACTGCTCCATTTTCGATAAAATTGTACCTTTAGGGTAATTAAGATT 2603_a12.seq									
8252		TCTTTCACTGCTCCATTTTCGATAAAATTGTACCTTTAGGGTAATTAAGATT 18rs21_a12.seq									
8160		TCTTTCACTGCTCCATTTTCGATAAAATTGTACCTTTAGGGTAATTAAGATT 515_a12.seq									
8149		TCTTTCACTGCTCCATTTTCGATAAAATTGTACCTTTAGGGTAATTAAGATT cjb111_a12.seq									
8202		TCTTTCACTGCTCCATTTTCGATAAAATTGTACCTTTAGGGTAATTAAGATT h36b_a12.seq									
		TAAATCTAAATAATGAAGTTTTTGTAAAGTTTCCAGAGATTATCTGTGTTT Majority									
		8360		8370		8380		8390		8400	
8302		TAAATCTAAATAATGAAGTTTTTGTAAAGTTTCCAGAGATTATCTGTGTTT 2603_a12.seq									
8302		TAAATCTAAATAATGAAGTTTTTGTAAAGTTTCCAGAGATTATCTGTGTTT 18rs21_a12.seq									
8210		TAAATCTAAATAATGAAGTTTTTGTAAAGTTTCCAGAGATTATCTGTGTTT 515_a12.seq									
8199		TAAATCTAAATAATGAAGTTTTTGTAAAGTTTCCAGAGATTATCTGTGTTT cjb111_a12.seq									
8252		TAAATCTAAATAATGAAGTTTTTGTAAAGTTTCCAGAGATTATCTGTGTTT h36b_a12.seq									
		GATAACTATCTAAGGGAAACAAAAGTAACTCTCCCCATTTCCCTTTTATA Majority									
		8410		8420		8430		8440		8450	
8352		GATAACTATCTAAGGGAAACAAAAGTAACTCTCCCCATTTCCCTTTTATA 2603_a12.seq									
8352		GATAACTATCTAAGGGAAACAAAAGTAACTCTCCCCATTTCCCTTTTATA 18rs21_a12.seq									
8260		GATAACTATCTAAGGGAAACAAAAGTAACTCTCCCCATTTCCCTTTTATA 515_a12.seq									
8249		GATAACTATCTAAGGGAAACAAAAGTAACTCTCCCCATTTCCCTTTTATA cjb111_a12.seq									
8302		GATAACTATCTAAGGGAAACAAAAGTAACTCTCCCCATTTCCCTTTTATA h36b_a12.seq									
		TCCTCGGGCTTATCAGTAAGTAGAAAATTACTTTTATTTAGATATCCATT Majority									
		8460		8470		8480		8490		8500	
8402		TCCTCGGGCTTATCAGTAAGTAGAAAATTACTTTTATTTAGATATCCATT 2603_a12.seq									
8402		TCCTCGGGCTTATCAGTAAGTAGAAAATTACTTTTATTTAGATATCCATT 18rs21_a12.seq									
8310		TCCTCGGGCTTATCAGTAAGTAGAAAATTACTTTTATTTAGATATCCATT 515_a12.seq									
8299		TCCTCGGGCTTATCAGTAAGTAGAAAATTACTTTTATTTAGATATCCATT cjb111_a12.seq									
8352		TCCTCGGGCTTATCAGTAAGTAGAAAATTACTTTTATTTAGATATCCATT h36b_a12.seq									
		TTTTTTTCATTTGTTCAAATTGGCTTTTCATATGATGCACCCAGTTTAAAAAT Majority									
		8510		8520		8530		8540		8550	
8452		TTTTTTTCATTTGTTCAAATTGGCTTTTCATATGATGCACCCAGTTTAAAAAT 2603_a12.seq									
8452		TTTTTTTCATTTGTTCAAATTGGCTTTTCATATGATGCACCCAGTTTAAAAAT 18rs21_a12.seq									
8360		TTTTTTTCATTTGTTCAAATTGGCTTTTCATATGATGCACCCAGTTTAAAAAT 515_a12.seq									
8349		TTTTTTTCATTTGTTCAAATTGGCTTTTCATATGATGCACCCAGTTTAAAAAT cjb111_a12.seq									
8402		TTTTTTTCATTTGTTCAAATTGGCTTTTCATATGATGCACCCAGTTTAAAAAT h36b_a12.seq									

FIGURE 19R



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T A T T A A T A G C A T A T G A T C T C G T A G G A A C A C C A T C A G T T A C A T G A A C A A T A Majority  
8560 8570 8580 8590 8600

8502 T A T T A A T A G C A T A T G A T C T C G T A G G A A C A C C A T C A G T T A C A T G A A C A A T A 2603\_al2.seq  
8502 T A T T A A T A G C A T A T G A T C T C G T A G G A A C A C C A T C A G T T A C A T G A A C A A T A 18rs21\_al2.seq  
8410 T A T T A A T A G C A T A T G A T C T C G T A G G A A C A C C A T C A G T T A C A T G A A C A A T A 515\_al2.seq  
8399 T A T T A A T A G C A T A T G A T C T C G T A G G A A C A C C A T C A G T T A C A T G A A C A A T A cjb111\_al2.seq  
8452 T A T T A A T A G C A T A T G A T C T C G T A G G A A C A C C A T C A G T T A C A T G A A C A A T A h36b\_al2.seq

A T T T T T T G A C T A T T T T C G A T T T A C T T G A C T C A A A A T A T C A T C T G C C T C C A T Majority  
8610 8620 8630 8640 8650

8552 A T T T T T T G A C T A T T T T C G A T T T A C T T G A C T C A A A A T A T C A T C T G C C T C C A T 2603\_al2.seq  
8552 A T T T T T T G A C T A T T T T C G A T T T A C T T G A C T C A A A A T A T C A T C T G C C T C C A T 18rs21\_al2.seq  
8460 A T T T T T T G A C T A T T T T C G A T T T A C T T G A C T C A A A A T A T C A T C T G C C T C C A T 515\_al2.seq  
8449 A T T T T T T G A C T A T T T T C G A T T T A C T T G A C T C A A A A T A T C A T C T G C C T C C A T cjb111\_al2.seq  
8502 A T T T T T T G A C T A T T T T C G A T T T A C T T G A C T C A A A A T A T C A T C T G C C T C C A T h36b\_al2.seq

G A A G G C T T T C A T A G T A A A T G T T T C T C C T A C T T T A C T A A G A T A G T A C T C C T Majority  
8660 8670 8680 8690 8700

8602 G A A G G C T T T C A T A G T A A A T G T T T C T C C T A C T T T A C T A A G A T A G T A C T C C T 2603\_al2.seq  
8602 G A A G G C T T T C A T A G T A A A T G T T T C T C C T A C T T T A C T A A G A T A G T A C T C C T 18rs21\_al2.seq  
8510 G A A G G C T T T C A T A G T A A A T G T T T C T C C T A C T T T A C T A A G A T A G T A C T C C T 515\_al2.seq  
8499 G A A G G C T T T C A T A G T A A A T G T T T C T C C T A C T T T A C T A A G A T A G T A C T C C T cjb111\_al2.seq  
8552 A A A A G C T T T C A T A G T A A A T G T T T C T C C T A C T T T A C T A A G A T A G T A C T C C T h36b\_al2.seq

T T T G T T G C T C T G G A G T T A A T C C A T T G G T A G T A G A T C C C C A C T T A G C T T T A Majority  
8710 8720 8730 8740 8750

8652 T T T G T T G C T C T G G A G T T A A T C C A T T G G T A G T A G A T C C C C A C T T A G C T T T A 2603\_al2.seq  
8652 T T T G T T G C T C T G G A G T T A A T C C A T T G G T A G T A G A T C C C C A C T T A G C T T T A 18rs21\_al2.seq  
8560 T T T G T T G C T C T G G A G T T A A T C C A T T G G T A G T A G A T C C C C A C T T A G C T T T A 515\_al2.seq  
8549 T T T G T T G C T C T G G A G T T A A T C C A T T G G T A G T A G A T C C C C A C T T A G C T T T A cjb111\_al2.seq  
8602 T T T G T T G C T C T G G A G T T A A T C C A T T G G T A G T A G A T C C C C A C T T A G C T T T A h36b\_al2.seq

G G A G C T T C T G T C G G A A T C C T T T T T A T A A T C T C T T C A G C A T T A T T T G T T A A Majority  
8760 8770 8780 8790 8800

8702 G G A G C T T C T G T C G G A A T C C T T T T T A T A A T C T C T T C A G C A T T A T T T G T T A A 2603\_al2.seq  
8702 G G A G C T T C T G T C G G A A T C C T T T T T A T A A T C T C T T C A G C A T T A T T T G T T A A 18rs21\_al2.seq  
8610 G G A G C T T C T G T C G G A A T C C T T T T T A T A A T C T C T T C A G C A T T A T T T G T T A A 515\_al2.seq  
8599 G G A G C T T C T G T C G G A A T C C T T T T T A T A A T C T C T T C A G C A T T A T T T G T T A A cjb111\_al2.seq  
8652 G G A G C T T C T T T A G G G A T C T T T T T A T A A T A T C A G C A G C A A T A T T A G T G A A h36b\_al2.seq

T T G T T T A T G A C T A T A A T T C T C T G T C T G A A T T G T G A A C T T A G T T T G A A G G C Majority  
8810 8820 8830 8840 8850

8752 T T G T T T A T G A C T A T A A T T C T C T G T C T G A A T T G T G A A C T T A G T T T G A A G G C 2603\_al2.seq  
8752 T T G T T T A T G A C T A T A A T T C T C T G T C T G A A T T G T G A A C T T A G T T T G A A G G C 18rs21\_al2.seq  
8660 T T G T T T A T G A C T A T A A T T C T C T G T C T G A A T T G T G A A C T T A G T T T G A A G G C 515\_al2.seq  
8649 T T G T T T A T G A C T A T A A T T C T C T G T C T G A A T T G T G A A C T T A G T T T G A A G G C cjb111\_al2.seq  
8702 T T T T T T A T A G C T A T A A T T A T T T G T C T G A A T T G T G A A C T T A G T T T T A A G G C h36b\_al2.seq

C A T A A T A T T T A T C A T C T T C T T T A A A T C C T T T T A C G A C A T C T A C A C T C C T A Majority  
8860 8870 8880 8890 8900

8802 C A T A A T A T T T A T C A T C T T C T T T A A A T C C T T T T A C G A C A T C T A C A C T C C T A 2603\_al2.seq  
8802 C A T A A T A T T T A T C A T C T T C T T T A A A T C C T T T T A C G A C A T C T A C A C T C C T A 18rs21\_al2.seq  
8710 C A T A A T A T T T A T C A T C T T C T T T A A A T C C T T T T A C G A C A T C T A C A C T C C T A 515\_al2.seq  
8699 C A T A A T A T T T A T C A T C T T C T T T A A A T C C T T T T A C G A C A T C T A C A C T C C T A cjb111\_al2.seq  
8752 C A T A A T A T T T A T C A T C T T C T T T A A A T C C T T T T A C G A C A T C T A C A C T C C T A h36b\_al2.seq

C C A T C A A A A A T A T C T G A A C C A T A G G T A A C T A A T G C A A C C C T A T T A T C A C T Majority  
8910 8920 8930 8940 8950

8852 C C A T C A A A A A T A T C T G A A C C A T A G G T A A C T A A T G C A A C C C T A T T A T C A C T 2603\_al2.seq  
8852 C C A T C A A A A A T A T C T G A A C C A T A G G T A A C T A A T G C A A C C C T A T T A T C A C T 18rs21\_al2.seq  
8760 C C A T C A A A A A T A T C T G A A C C A T A G G T A A C T A A T G C A A C C C T A T T A T C A C T 515\_al2.seq  
8749 C C A T C A A A A A T A T C T G A A C C A T A G G T A A C T A A T G C A A C C C T A T T A T C A C T cjb111\_al2.seq  
8799 C C A T C A A A A A T A T C T G A A C C A T A G T A A C T A A G C T G C T C G G T T T C A A C h36b\_al2.seq

G T T T G C T C C T A A A A T A T C T T T T A C T G C G G T C C C A A G A G C T T C G G C A G C T T Majority  
8960 8970 8980 8990 9000

8902 G T T T G C T C C T A A A A T A T C T T T T A C T G C G G T C C C A A G A G C T T C G G C A G C T T 2603\_al2.seq  
8902 G T T T G C T C C T A A A A T A T C T T T T A C T G C G G T C C C A A G A G C T T C G G C A G C T T 18rs21\_al2.seq  
8810 G T T T G C T C C T A A A A T A T C T T T T A C T G C G G T C C C A A G A G C T T C G G C A G C T T 515\_al2.seq  
8799 G T T T G C T C C T A A A A T A T C T T T T A C T G C G G T C C C A A G A G C T T C G G C A G C T T cjb111\_al2.seq  
8849 A T T T G C T C C T A A A A T A T C T T T T A T A A T T G T T T C T A A G T G C T T C A C C T G C C T h36b\_al2.seq

FIGURE 19S

**FIGURE 19T**



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T C T T T T T A T A A C C T T C G G G C G C T G T T T C T T C T G A T A A A G T A T A A T C T C C A Majority									
9460		9470		9480		9490		9500	
9402	T	C	T	T	T	T	A	T	A
9402	T	C	T	T	T	T	A	T	A
9310	T	C	T	T	T	T	A	T	A
9299	T	C	T	T	T	T	A	T	A
9337	T	C	T	T	T	T	A	T	A
C G T A T G A G A T T A T C A A A A G T A G C T T C A C C T G T T A G C T C A G C A G T T A C T T T Majority									
9510		9520		9530		9540		9550	
9452	C	G	T	A	T	G	A	G	A
9452	C	G	T	A	T	G	A	G	A
9360	C	G	T	A	T	G	A	G	A
9349	C	G	T	A	T	G	A	G	A
9387	C	G	T	A	T	G	A	G	A
T T C T A T T T T A C T T T C T G G A T G A G C A G T A G T T T T T A A A A C A A A G G T A G C T T Majority									
9560		9570		9580		9590		9600	
9502	T	T	C	T	A	T	T	T	A
9502	T	T	C	T	A	T	T	T	A
9410	T	T	C	T	A	T	T	T	A
9399	T	T	C	T	A	T	T	T	A
9437	T	T	C	T	A	T	T	T	A
T T G A A A G T G G T T T G T T C T G G T C A T C T G T C T T T T T A A C A A C T A A C T T T C C T Majority									
9610		9620		9630		9640		9650	
9552	T	T	G	A	A	G	T	G	G
9552	T	T	G	A	A	G	T	G	G
9460	T	T	G	A	A	G	T	G	G
9449	T	T	G	A	A	G	T	G	G
9487	T	T	G	A	A	G	T	G	G
T T A G C A C C A T T T T C C G G T A C G G T A C T T T C C C C T A A A A C A T T G G T A T T A A G Majority									
9660		9670		9680		9690		9700	
9602	T	T	A	G	C	A	C	C	A
9602	T	T	A	G	C	A	C	C	A
9510	T	T	A	G	C	A	C	C	A
9499	T	T	A	G	C	A	C	C	A
9537	T	T	A	G	C	A	C	C	A
C G G T A T T T G C G A C A A A C A A A A A A G A C T T A A C G T C A A T A T T T T A G A A A A T T Majority									
9710		9720		9730		9740		9750	
9652	C	G	G	T	A	T	T	G	C
9652	C	G	G	T	A	T	T	G	C
9560	C	G	G	T	A	T	T	G	C
9549	C	G	G	T	A	T	T	G	C
9587	C	G	G	T	A	T	T	G	C
T T T G G T A T T T T C T C A T T T T A C A A C T C C T A T T G T G C C G A A A T G T C G T T T C T Majority									
9760		9770		9780		9790		9800	
9702	T	T	T	G	G	T	A	T	T
9702	T	T	T	G	G	T	A	T	T
9610	T	T	T	G	G	T	A	T	T
9599	T	T	T	G	G	T	A	T	T
9637	T	T	T	G	G	T	A	T	T
A A A T C T A A G A T C A G A T A C A G A A T A T C C T A G A A T A T A C A A A C T A T C A C T T A Majority									
9810		9820		9830		9840		9850	
9752	A	A	A	T	C	T	A	A	G
9752	A	A	A	T	C	T	A	A	G
9660	A	A	A	T	C	T	A	A	G
9649	A	A	A	T	C	T	A	A	G
9687	A	A	A	T	C	T	A	A	G
T T A T G A T A T C A A T A A T T T C T T A T T A T A A G G T A T G G A A T T T T A A T G T T T T T Majority									
9860		9870		9880		9890		9900	
9802	T	T	A	T	G	A	T	A	T
9802	T	T	A	T	G	A	T	A	T
9710	T	T	A	T	G	A	T	A	T
9699	T	T	A	T	G	A	T	A	T
9737	T	T	A	T	G	A	T	A	T

FIGURE 19U

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	T C C C A A T T T T T G A A T A A T T T T T C T T T T T A T T T G A T A A T C T T A T T T T T T A T	Majority
	9910 9920 9930 9940 9950	
9852	T C C C A A T T T T T G A A T A A T T T T T C T T T T T A T T T G A T A A T C T T A T T T T T T A T	2603_ai2.seq
9852	T C C C A A T T T T T G A A T A A T T T T T C T T T T T A T T T G A T A A T C T T A T T T T T T A T	18rs21_ai2.seq
9760	T C C C A A T T T T T G A A T A A T T T T T C T T T T T A T T T G A T A A T C T T A T T T T T T A T	515_ai2.seq
9749	T C C C A A T T T T T G A A T A A T T T T T C T T T T T A T T T G A T A A T C T T A T T T T T T A T	cjb111_ai2.seq
9787	T C C C A A T T T T T G A A T A A T T T T T C T T T T T A T T T G A T A A T C T T A T T T T T T A T	h36b_ai2.seq
	T A T C T T A G A A A T A T T T T C A A T G A G C T T A A G T A G T T G A T T T T T C T T T T T T T A	Majority
	9960 9970 9980 9990 10000	
9902	T A T C T T A G A A A T A T T T T C A A T G A G C T T A A G T A G T T G A T T T T T C T T T T T T T A	2603_ai2.seq
9902	T A T C T T A G A A A T A T T T T C A A T G A G C T T A A G T A G T T G A T T T T T C T T T T T T T A	18rs21_ai2.seq
9808	T A T C T T A G A A A T A T T T T C A A T G A G C T T A A G T A G T T G A T T T T T C T T T T T T T A	515_ai2.seq
9799	T A T C T T A G A A A T A T T T T C A A T G A G C T T A A G T A G T T G A T T T T T C T T T T T T T A	cjb111_ai2.seq
9837	T A T C T T A G A A A T A T T T T C A A T G A G C T T A A G T A G T T G A T T T T T C T T T T T T T A	h36b_ai2.seq
	T G T T T T A A A A T A T T T G C T T A A A A A T A A T G T T T G A G A G A G - - T T T A C T G A A T	Majority
	10010 10020 10030 10040 10050	
9952	T G T T T T A A A A T A T T T G C T T A A A A A T A A T G T T T G A G A G A G - - T T T A C T G A A T	2603_ai2.seq
9952	T G T T T T A A A A T A T T T G C T T A A A A A T A A T G T T T G A G A G A G - - T T T A C T G A A T	18rs21_ai2.seq
9856	T G T T T T A A A A T A T T T G C T T A A A A A T A A T G T T T G A G A G A G - - T T T A C T G A A T	515_ai2.seq
9849	T G T T T T A A A A T A T T T G C T T A A A A A T A A T G T T T G A G A G A G - - T T T A C T G A A T	cjb111_ai2.seq
9887	T G T T T T A A A A T A T T T G C T T A A A A A T A A T G T T T G A G A G A G - - T T T A C T G A A T	h36b_ai2.seq
	T G A T T G A A A A T T A T T T T A G A A A A A G A C A T C C T T A A T C A A A T A A A A C T T C T A	Majority
	10060 10070 10080 10090 10100	
10000	T G A T T G A A A A T T A T T T T A G A A A A A G A C A T C C T T A A T C A A A T A A A A C T T C T A	2603_ai2.seq
10000	T G A T T G A A A A T T A T T T T A G A A A A A G A C A T C C T T A A T C A A A T A A A A C T T C T A	18rs21_ai2.seq
9903	T G A T T G A A A A T T A T T T T A G A A A A A G A C A T C C T T A A T C A A A T A A A A C T T C T A	515_ai2.seq
9897	T G A T T G A A A A T T A T T T T A G A A A A A G A C A T C C T T A A T C A A A T A A A A C T T C T A	cjb111_ai2.seq
9937	T G A T T G A A A A T T A T T T T A G A A A A A G A C A T C C T T A A T C A A A T A A A A C T T C T A	h36b_ai2.seq
	A C T T T A T G C T A T G A T T A C T A C C C T T C C A T T A C T C T A G A C A A A T C A T G T C A	Majority
	10110 10120 10130 10140 10150	
10050	A C T T T A T G C T A T G A T T A C T A C C C T T C C A T T A C T C T A G A C A A A T C A T G T C A	2603_ai2.seq
10050	A C T T T A T G C T A T G A T T A C T A C C C T T C C A T T A C T C T A G A C A A A T C A T G T C A	18rs21_ai2.seq
9953	A C T T T A T G C T A T G A T T A C T A C C C T T C C A T T A C T C T A G A C A A A T C A T G T C A	515_ai2.seq
9947	A C T T T A T G C T A T G A T T A C T A C C C T T C C A T T A C T C T A G A C A A A T C A T G T C A	cjb111_ai2.seq
9987	A C T T T A T G C T A T G A T T A C T A C C C T T C C A T T A C T C T A G A C A A A T C A T G T C A	h36b_ai2.seq
	T C A A C T T G G T T T A T C T G A A C T A C T T A T T A G G A A - A T A T T G T C A T G A T T T A	Majority
	10160 10170 10180 10190 10200	
10100	T C A A C T T G G T T T A T C T G A A C T A C T T A T T A G G A A - A T A T T G T C A T G A T T T A	2603_ai2.seq
10100	T C A A C T T G G T T T A T C T G A A C T A C T T A T T A G G A A - A T A T T G T C A T G A T T T A	18rs21_ai2.seq
10003	T C A A C T T G G T T T A T C T G A A C T A C T T A T T A G G A A - A T A T T G T C A T G A T T T A	515_ai2.seq
9997	T C A A C T T G G T T T A T C T G A A C T A C T T A T T A G G A A - A T A T T G T C A T G A T T T A	cjb111_ai2.seq
10037	T C A A C T T G G T T T A T C T G A A C T A C T T A T T A G G A A - A T A T T G T C A T G A T T T A	h36b_ai2.seq
	A C A A C T T T A T T T T A A C A G T C A A C T C T C T C T G A A T A T C G A A A A G A G T A C A A T	Majority
	10210 10220 10230 10240 10250	
10149	A C A A C T T T A T T T T A A C A G T C A A C T C T C T C T G A A T A T C G A A A A G A G T A C A A T	2603_ai2.seq
10149	A C A A C T T T A T T T T A A C A G T C A A C T C T C T C T G A A T A T C G A A A A G A G T A C A A T	18rs21_ai2.seq
10053	A C A A C T T T A T T T T A A C A G T C A A C T C T C T C T G A A T A T C G A A A A G A G T A C A A T	515_ai2.seq
10046	A C A A C T T T A T T T T A A C A G T C A A C T C T C T C T G A A T A T C G A A A A G A G T A C A A T	cjb111_ai2.seq
10086	A C A A C T T T A T T T T A A C A G T C A A C T C T C T C T G A A T A T C G A A A A G A G T A C A A T	h36b_ai2.seq
	A G T C T A C C A A T C T A A T G G T G T A A C T A G A G A A C A A G C T T T C A A A T A T A T T T	Majority
	10260 10270 10280 10290 10300	
10199	A G T C T A C C A A T C T A A T G G T G T A A C T A G A G A A C A A G C T T T C A A A T A T A T T T	2603_ai2.seq
10199	A G T C T A C C A A T C T A A T G G T G T A A C T A G A G A A C A A G C T T T C A A A T A T A T T T	18rs21_ai2.seq
10103	A G T C T A C C A A T C T A A T G G T G T A A C T A G A G A A C A A G C T T T C A A A T A T A T T T	515_ai2.seq
10096	A G T C T A C C A A T C T A A T G G T G T A A C T A G A G A A C A A G C T T T C A A A T A T A T T T	cjb111_ai2.seq
10136	A G T C T A C C A A T C T A A T G G T G T A A C T A G A G A A C A A G C T T T C A A A T A T A T T T	h36b_ai2.seq
	A T C A T C A A T C A C A C G T T T T A C A A C T T T T A A A A T T T T T G A T C A C G A A T G A T	Majority
	10310 10320 10330 10340 10350	
10249	A T C A T C A A T C A C A C G T T T T A C A A C T T T T A A A A T T T T T G A T C A C G A A T G A T	2603_ai2.seq
10249	A T C A T C A A T C A C A C G T T T T A C A A C T T T T A A A A T T T T T G A T C A C G A A T G A T	18rs21_ai2.seq
10152	A T C A T C A A T C A C A C G T T T T A C A A C T T T T A A A A T T T T T G A T C A C G A A T G A T	515_ai2.seq
10146	A T C A T C A A T C A C A C G T T T T A C A A C T T T T A A A A T T T T T G A T C A C G A A T G A T	cjb111_ai2.seq
10186	A T C A T C A A T C A C A C G T T T T A C A A C T T T T A A A A T T T T T G A T C A C G A A T G A T	h36b_ai2.seq

FIGURE 19V



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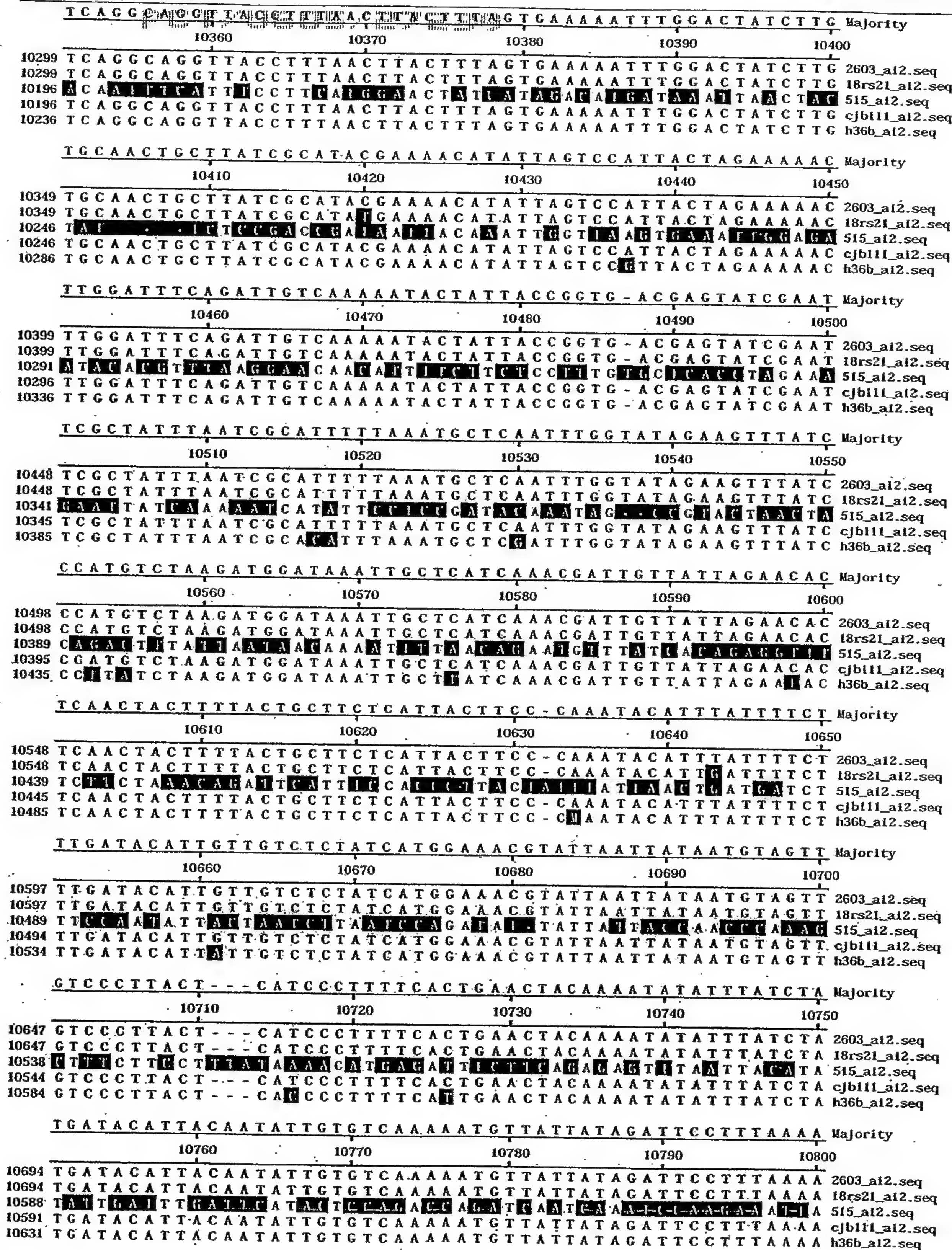


FIGURE 19W

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TTAATTTTAAAGAGAGATATAGACTATATTTTCTTGC-----T Majority									
10810		10820		10830		10840		10850	
10744	TTAATTTTAAAGAGAGATATAGACTATATTTTCTTGC-----T	2603_al2.seq							
10744	TTAATTTTAAAGAGAGATATAGACTATATTTTCTTGC-----T	18rs21_al2.seq							
10638	TTTCAATCAATACAGAGAGAAATATTTTCTTGC-----T	515_al2.seq							
10641	TTAATTTTAAAGAGAGATATAGACTATATTTTCTTGC-----T	cjb111_al2.seq							
10681	TTAATTTTAAAGAGAGATATAGACTATATTTTCTTGC-----T	h36b_al2.seq							
TACCTTACTTCCCATAACTCTTTTCTAATCCAAATTGGAC--TGAGAAAG Majority									
10860		10870		10880		10890		10900	
10785	TACCTTACTTCCCATAACTCTTTTCTAATCCAAATTGGAC--TGAGAAAG	2603_al2.seq							
10785	TACCTTACTTCCCATAACTCTTTTCTAATCCAAATTGGAC--TGAGAAAG	18rs21_al2.seq							
10688	GAATAAGTACCCGTAGCTCTTAATAACCTTAAATAATTAACATTAAGAG	515_al2.seq							
10682	TACCTTACTTCCCATAACTCTTTTCTAATCCAAATTGGAC--TGAGAAAG	cjb111_al2.seq							
10722	TACCTTACTTCCCATAACTCTTTTCTAATCCAAATTGGAC--TGAGAAAG	h36b_al2.seq							
CGTATCGATAATGTAATA-GCTATTTTTCGAAAATTATCCCAAATTCCAAA Majority									
10910		10920		10930		10940		10950	
10833	CGTATCGATAATGTAATA-GCTATTTTTCGAAAATTATCCCAAATTCCAAA	2603_al2.seq							
10833	CGTATCGATAATGTAATA-GCTATTTTTCGAAAATTATCCCAAATTCCAAA	18rs21_al2.seq							
10738	CTAGAGGATTAACAAATGCTCTAATTTTAAATGTTAAATTTTTTGAATA	515_al2.seq							
10730	CGTATCGATAATGTAATA-GCTATTTTTCGAAAATTATCCCAAATTCCAAA	cjb111_al2.seq							
10770	CGTATCGATAATGTAATA-GCTATTTTTCGAAAATTATCCCAAATTCCAAA	h36b_al2.seq							
AATTATTACAGCCCACTCAAAGATGCTCTTCC--CTTATCTGGCTCCTATC Majority									
10960		10970		10980		10990		11000	
10882	AATTATTACAGCCCACTCAAAGATGCTCTTCC--CTTATCTGGCTCCTATC	2603_al2.seq							
10882	AATTATTACAGCCCACTCAAAGATGCTCTTCC--CTTATCTGGCTCCTATC	18rs21_al2.seq							
10788	ATATAATCAACCTTTCAACCTGTTTTCCTAGGTTGAATGTTCTTTTAAAT	515_al2.seq							
10779	AATTATTACAGCCCACTCAAAGATGCTCTTCC--CTTATCTGGCTCCTATC	cjb111_al2.seq							
10819	AATTATTACAGCCCACTCAAAGATGCTCTTCC--CTTATCTGGCTCCTATC	h36b_al2.seq							
ATGATGAGTTGGTAAAAGTTGCTATCTTTTTTTTCCGAACATTTATTTTA Majority									
11010		11020		11030		11040		11050	
10930	ATGATGAGTTGGTAAAAGTTGCTATCTTTTTTTTCCGAACATTTATTTTA	2603_al2.seq							
10930	ATGATGAGTTGGTAAAAGTTGCTATCTTTTTTTTCCGAACATTTATTTTA	18rs21_al2.seq							
10838	TCCTTACGCAAAATTCCTGTTGCTAGTTTCTGCTCTTAAAGGCTTATCTTTTA	515_al2.seq							
10827	ATGATGAGTTGGTAAAAGTTGCTATCTTTTTTTTCCGAACATTTATTTTA	cjb111_al2.seq							
10867	ATGATGAGTTGGTAAAAGTTGCTATCTTTTTTTTCCGAACATTTATTTTA	h36b_al2.seq							
GGATTAAATCAATTAAATCCCTGAAACAATT--TCATTTCTTTCATGGAAC Majority									
11060		11070		11080		11090		11100	
10979	GGATTAAATCAATTAAATCCCTGAAACAATT--TCATTTCTTTCATGGAAC	2603_al2.seq							
10980	GGATTAAATCAATTAAATCCCTGAAACAATT--TCATTTCTTTCATGGAAC	18rs21_al2.seq							
10888	CTAATAAATCAAGAGATCTATGAGAGTGAAGTATTTTCTTCTCAAGAG	515_al2.seq							
10877	GGATTAAATCAATTAAATCCCTGAAACAATT--TCATTTCTTTCATGGAAC	cjb111_al2.seq							
10917	GGATTAAATCAATTAAATCCCTGAAACAATT--TCATTTCTTTCATGGAAC	h36b_al2.seq							
TATCATAGACATGATAAATTAACACTACTATTCTCCGACCGATAATTACAAA Majority									
11110		11120		11130		11140		11150	
11027	TATCATAGACATGATAAATTAACACTACTATTCTCCGACCGATAATTACAAA	2603_al2.seq							
11028	TATCATAGACATGATAAATTAACACTACTATTCTCCGACCGATAATTACAAA	18rs21_al2.seq							
10938	ATCTCTAAGTCAAGGCTATCTGAATACTTTAACTCTGCTGAGCGGCACTTC	515_al2.seq							
10925	TATCATAGACATGATAAATTAACACTACTATTCTCCGACCGATAATTACAAA	cjb111_al2.seq							
10965	TATCATAGACATGATAAATTAACACTACTATTCTCCGACCGATAATTACAAA	h36b_al2.seq							
TTGGTTAAGTAAAATTGGAGAATA--CACGTTTAAAGAACAACATTTTTC Majority									
11160		11170		11180		11190		11200	
11077	TTGGTTAAGTAAAATTGGAGAATA--CACGTTTAAAGAACAACATTTTTC	2603_al2.seq							
11078	TTGGTTAAGTAAAATTGGAGAATA--CACGTTTAAAGAACAACATTTTTC	18rs21_al2.seq							
10987	TTGCAATATATAGGATTTGAAGAGTAGACCAAGTTCTTAAATAAGAGGTTGGT	515_al2.seq							
10975	TTGGTTAAGTAAAATTGGAGAATA--CACGTTTAAAGAACAACATTTTTC	cjb111_al2.seq							
11015	TTGGTTAAGTAAAATTGGAGAATA--CACGTTTAAAGAACAACATTTTTC	h36b_al2.seq							
TTCTCCTTTTGTACTCATCTAGAAAGAAATCATCAAAAATCATATTCTCTCCG Majority									
11210		11220		11230		11240		11250	
11124	TTCTCCTTTTGTACTCATCTAGAAAGAAATCATCAAAAATCATATTCTCTCCG	2603_al2.seq							
11125	TTCTCCTTTTGTACTCATCTAGAAAGAAATCATCAAAAATCATATTCTCTCCG	18rs21_al2.seq							
11037	AAATCTCTGTAATCATCTAGAAAGAAATCATCAAAAATCATATTCTCTCCG	515_al2.seq							
11022	TTCTCCTTTTGTACTCATCTAGAAAGAAATCATCAAAAATCATATTCTCTCCG	cjb111_al2.seq							
11062	TTCTCCTTTTGTACTCATCTAGAAAGAAATCATCAAAAATCATATTCTCTCCG	h36b_al2.seq							

FIGURE 19X



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A T A C A G A - - T A G C C G T A C T A A C T A C A G A C T T T A T T A A T A A C C A A A T T T T A Majority									
11260		11270		11280		11290		11300	
11174	A T A C A G A - -	T A G C C G T A C T A A C T A C A G A C T T T A T T A A T A A C C A A A T T T T A	2603_al2.seq						
11175	A T A C A G A - -	T A G C C G T A C T A A C T A C A G A C T T T A T T A A T A A C C A A A T T T T A	18rs21_al2.seq						
11086	I T A A A G A C A T A G T C T G T T C A A A G I I I A A I T T C C C A A A A A G I T A A T G T G T	515_al2.seq							
11072	A T A C A G A - -	T A G C C G T A C T A A C T A C A G A C T T T A T T A A T A A C C A A A T T T T A	cjb111_al2.seq						
11112	A T A C A G A - -	T A G C C G T A C T A A C T A C A G A C T T T A T T A A T A A C C A A A T T T T A	h36b_al2.seq						
A C A G A A T G C T T A T T A C A G A G A T T T T C T T C T A A A A A G A T T C A T T T C C A C C C Majority									
11310		11320		11330		11340		11350	
11222	A C A G A A T G C T T A T T A C A G A G A T T T T C T T C T A A A A A G A T T C A T T T C C A C C C	2603_al2.seq							
11223	A C A G A A T G C T T A T T A C A G A G A T T T T C T T C T A A A A A G A T T C A T T T C C A C C C	18rs21_al2.seq							
11136	I T G G A G T G A T A T T T T C I . . . . . C T T T C A A A T G I G G I A A T T C A G G T C C G I C T	515_al2.seq							
11120	A C A G A A T G C T T A T T A C A G A G A T T T T C T T C T A A A A A G A T T C A T T T C C A C C C	cjb111_al2.seq							
11160	A C A G A A T G C T T A T T A C A G A G A T T T T C T T C T A A A A A G A T T C A T T T C C A C C C	h36b_al2.seq							
T T A C T A T T T A T T A A C T G A T G A T C T T T C C A A T A T T A C T A A T C T T A A T C C A G Majority									
11360		11370		11380		11390		11400	
11272	T T A C T A T T T A T T A A C T G A T G A T C T T T C C A A T A T T A C T A A T C T T A A T C C A G	2603_al2.seq							
11273	T T A C T A T T T A T T A A C T G A T G A T C T T T C C A A T A T T A C T A A T C T T A A T C C A G	18rs21_al2.seq							
11181	C C T G C A A T C T G T A A A T A A A G A T T T T T A G A G T A G T G . T G A A A T G A A . . . A A	515_al2.seq							
11170	T T A C T A T T T A T T A A C T G A T G A T C T T T C C A A T A T T A C T A A T C T T A A T C C A G	cjb111_al2.seq							
11210	T T A C T A T T T A T T A A C T G A T G A T C T T T C C A A T A T T A C T A A T C T T A A T C C A G	h36b_al2.seq							
A C A T T A T T A T T A C C A A T A A A A A G C T T T T C T C C T T T A T C A A A C A T G A G A T T Majority									
11410		11420		11430		11440		11450	
11322	A C A T T A T T A T T A C C A A T A A A A A G C T T T T C T C C T T T A T C A A A C A T G A G A T T	2603_al2.seq							
11323	A C A T T A T T A T T A C C A A T A A A A A G C T T T T C T C C T T T A T C A A A C A T G A G A T T	18rs21_al2.seq							
11227	A T G G T T C T A A G A G C A A T T A A A I G C T T T T T C I T T A A T . . . . . A A T T C	515_al2.seq							
11220	A C A T T A T T A T T A C C A A T A A A A A G C T T T T C T C C T T T A T C A A A C A T G A G A T T	cjb111_al2.seq							
11260	A C A T T A T T A T T A C C A A T A A A A A G C T T T T C T C C T T T A T C A A A C A T G A G A T T	h36b_al2.seq							
T C T T C A G A G A G T T T A A T T A C A T A T A T T G A T C T G A T T A A T A C T T C A G A C C A Majority									
11460		11470		11480		11490		11500	
11372	T C T T C A G A G A G T T T A A T T A C A T A T A T T G A T C T G A T T A A T A C T T C A G A C C A	2603_al2.seq							
11373	T C T T C A G A G A G T T T A A T T A C A T A T A T T G A T C T G A T T A A T A C T T C A G A C C A	18rs21_al2.seq							
11269	T A C C A G C A T A A G T G A I G A A A A T A T C A T A G C A G A T T I T T C A A G G I A A G C T	515_al2.seq							
11270	T C T T C A G A G A G T T T A A T T A C A T A T A T T G A T C T G A T T A A T A C T T C A G A C C A	cjb111_al2.seq							
11310	T C T T C A G A G A G T T T A A T T A C A T A T A T T G A T C T G A T T A A T A C T T C A G A C C A	h36b_al2.seq							
A A T C A A T C A A A T C C A A A A A A T T A T T T C A T C A A T A C A G G A A G A A A A A T A T T Majority									
11510		11520		11530		11540		11550	
11422	A A T C A A T C A A A T C C A A A A A A T T A T T T C A T C A A T A C A G G A A G A A A A A T A T T	2603_al2.seq							
11423	A A T C A A T C A A A T C C A A A A A A T T A T T T C A T C A A T A C A G G A A G A A A A A T A T T	18rs21_al2.seq							
11319	G T G T C A G C A A A A T C A G A G C T A G A T T I C A G A T A C G G A A T T A I A A A T A A C	515_al2.seq							
11320	A A T C A A T C A A A T C C A A A A A A T T A T T T C A T C A A T A C A G G A A G A A A A A T A T T	cjb111_al2.seq							
11360	A A T C A A T C A A A T C C A A A A A A T T A T T T C A T C A A T A C A G G A A G A A A A A T A T T	h36b_al2.seq							
G C A A A C T T T T T T G C A A A A A C T A A T G A A A T A A C T A A T C G T A G C T C C T - - - - Majority									
11560		11570		11580		11590		11600	
11472	G C A A A C T T T T T T G C A A A A A C T A A T G A A A T A A C T A A T C G T A G C T C C T - - - -	2603_al2.seq							
11473	G C A A A C T T T T T T G C A A A A A C T A A T G A A A T A A C T A A T C G T A G C T C C T - - - -	18rs21_al2.seq							
11369	I C C I T I A G C T T C T A I A T I A A A A T G I I T A A C G A I T C A A G C T I C T C I G G	515_al2.seq							
11370	G C A A A C T T T T T T G C A A A A A C T A A T G A A A T A A C T A A T C G T A G C T C C T - - - -	cjb111_al2.seq							
11410	G C A A A C T T T T T T G C A A A A A C T A A T G A A A T A A C T A A T C G T A G C T C C T - - - -	h36b_al2.seq							
A T A A C T C T T A A A A A T T A A C A T T A A A A A G C T - A G A G C A T T G T G T A A T G C T C Majority									
11610		11620		11630		11640		11650	
11517	A T A A C T C T T A A A A A T T A A C A T T A A A A A G C T - A G A G C A T T G T G T A A T G C T C	2603_al2.seq							
11518	A T A A C T C T T A A A A A T T A A C A T T A A A A A G C T - A G A G C A T T G T G T A A T G C T C	18rs21_al2.seq							
11419	A T A C G C A T A A A A A T T T G G A C C A T A A T G C T T A A C A C G C G C T G T G A G A . . .	515_al2.seq							
11415	A T A A C T C T T A A A A A T T A A C A T T A A A A A G C T - A G A G C A T T G T G T A A T G C T C	cjb111_al2.seq							
11455	A T A A C T C T T A A A A A T T A A C A T T A A A A A G C T - A G A G C A T T G T G T A A T G C T C	h36b_al2.seq							
T A G C T T T T T A A T G T T A A T T T T T T T G A A T A A T A A T C C A A C T T T T C A A C T Majority									
11660		11670		11680		11690		11700	
11566	T A G C T T T T T A A T G T T A A T T T T T T T T G A A T A A T A A T C C A A C T T T T C A A C T	2603_al2.seq							
11567	T A G C T T T T T A A T G T T A A T T T T T T T T G A A T A A T A A T C C A A C T T T T C A A C T	18rs21_al2.seq							
11466	A G A T G T T T C A T A G A T A G C T C A A A C A A A A T G T A A A A A C G A . T T A T G A C A	515_al2.seq							
11464	T A G C T T T T T A A T G T T A A T T T T T T T T G A A T A A T A A T C C A A C T T T T C A A C T	cjb111_al2.seq							
11504	T A G C T T T T T A A T G T T A A T T T T T T T T G A A T A A T A A T C C A A C T T T T C A A C T	h36b_al2.seq							

FIGURE 19Y



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G T T T T T T C C C A T G T G A A A T G T T C T T T A A T T C T T T T A G C A A T A T T C T G T T G Majority									
11710		11720		11730		11740		11750	
11616	G T T T T T T C C C A T G T G A A A T G T T C T T T A A T T C T T T T A G C A A T A T T C T G T T G	2603_al2.seq							
11617	G T T T T T T C C C A T G T G A A A T G T T C T T T A A T T C T T T T A G C A A T A T T C T G T T G	18rs21_al2.seq							
11514	G A A A A A T G A C I T G A C C C A T G G T C T A A A A A A T A C T A G C I A A A T G G T G T T	515_al2.seq							
11514	G T T T T T T C C C A T G T G A A A T G T T C T T T A A T T C T T T T A G C A A T A T T C T G T T G	cjb111_al2.seq							
11554	G T T T T T T C C C A T G T G A A A T G T T C T T T A A T T C T T T T A G C A A T A T T C T G T T G	h36b_al2.seq							
T A G T T T C T C T C T T A A T G C C T T A T C T T T T A C T A A T A A A T C A A G A G A T T C A T Majority									
11760		11770		11780		11790		11800	
11666	T A G T T T C T C T C T T A A T G C C T T A T C T T T T A C T A A T A A A T C A A G A G A T T C A T	2603_al2.seq							
11667	T A G T T T C T C T C T T A A T G C C T T A T C T T T T A C T A A T A A A T C A A G A G A T T C A T	18rs21_al2.seq							
11564	C P T T T G C A A A A G A T A G C C C I T C I A A C G T T G T T A A C I G A A A A C G C G I A T I A C	515_al2.seq							
11564	T A G T T T C T C T C T T A A T G C C T T A T C T T T T A C T A A T A A A T C A A G A G A T T C A T	cjb111_al2.seq							
11604	T A G T T T C T C T C T T A A T G C C T T A T C T T T T A C T A A T A A A T C A A G A G A T T C A T	h36b_al2.seq							
G G A G T - - - G A C T G A G T A T T T T C T T C C A T G A T G A T T C C T A A C T C A G G G C T A Majority									
11810		11820		11830		11840		11850	
11716	G G A G T - - - G A C T G A G T A T T T T C T T C C A T G A T G A T T C C T A A C T C A G G G C T A	2603_al2.seq							
11717	G G A G T - - - G A C T G A G T A T T T T C T T C C A T G A T G A T T C C T A A C T C A G G G C T A	18rs21_al2.seq							
11614	A A A I C A C A A A A T C A A T A T T T T C A T C I G A A A C A T A T T T A I C A G C G I C I T G	515_al2.seq							
11614	G G A G T - - - G A C T G A G T A T T T T C T T C C A T G A T G A T T C C T A A C T C A G G G C T A	cjb111_al2.seq							
11654	G G A G T - - - G A C T G A G T A T T T T C T T C C A T G A T G A T T C C T A A C T C A G G G C T A	h36b_al2.seq							
T - - - - - C A A T A A C T T C A A C T G T T C C A C C G C G A T C T G T T G C A A T A A T A G C Majority									
11860		11870		11880		11890		11900	
11763	T - - - - - C A A T A A C T T C A A C T G T T C C A C C G C G A T C T G T T G C A A T A A T A G C	2603_al2.seq							
11764	T - - - - - C A A T A A C T T C A A C T G T T C C A C C G C G A T C T G T T G C A A T A A T A G C	18rs21_al2.seq							
11664	T A T T C T C A I T T T T G T I A A I A A T A G G A I A G C G T G C T T G A C A A T A T T T T I	515_al2.seq							
11661	T - - - - - C A A T A A C T T C A A C T G T T C C A C C G C G A T C T G T T G C A A T A A T A G C	cjb111_al2.seq							
11701	T - - - - - C A A T A A C T T C A A C T G T T C C A C C G C G A T C T G T T G C A A T A A T A G C	h36b_al2.seq							
A C T T G A A A G T A G A C C A G C T T C T A A A - - - - A T A G A G G T T G G T A A T C C C T C Majority									
11910		11920		11930		11940		11950	
11807	A C T T G A A A G T A G A C C A G C T T C T A A A - - - - A T A G A G G T T G G T A A T C C C T C	2603_al2.seq							
11808	A C T T G A A A G T A G A C C A G C T T C T A A A - - - - A T A G A G G T T G G T A A T C C C T C	18rs21_al2.seq							
11714	G G T G G G I A A A C G G I A A A T T T I T C I A C C C I I G T C P I C A T I I A T A A T C G G T	515_al2.seq							
11705	A C T T G A A A G T A G A C C A G C T T C T A A A - - - - A T A G A G G T T G G T A A T C C C T C	cjb111_al2.seq							
11745	A C T T G A A A G T A G A C C A G C T T C T A A A - - - - A T A G A G G T T G G T A A T C C C T C	h36b_al2.seq							
T G G A T A C A T T G A A G G G T A A A C A A A G A T A T C A G T C T G T - G C C A T T A A A G A C Majority									
11960		11970		11980		11990		12000	
11852	T G G A T A C A T T G A A G G G T A A A C A A A G A T A T C A G T C T G T - G C C A T T A A A G A C	2603_al2.seq							
11853	T G G A T A C A T T G A A G G G T A A A C A A A G A T A T C A G T C T G T - G C C A T T A A A G A C	18rs21_al2.seq							
11763	A A A I C A C A T G A T I A G T I G T T A C A A P A A C A A C A C G G T A G C C A C G G T I A A C	515_al2.seq							
11750	T G G A T A C A T T G A A G G G T A A A C A A A G A T A T C A G T C T G T - G C C A T T A A A G A C	cjb111_al2.seq							
11790	T G G A T A C A T T G A A G G G T A A A C A A A G A T A T C A G T C T G T - G C C A T T A A A G A C	h36b_al2.seq							
A T A G T C T G T T C A A A G T T T A A T T T C C C C A A A A A G T T A A T C T G T T T G G A C T G Majority									
12010		12020		12030		12040		12050	
11901	A T A G T C T G T T C A A A G T T T A A T T T C C C C A A A A A G T T A A T C T G T T T G G A C T G	2603_al2.seq							
11902	A T A G T C T G T T C A A A G T T T A A T T T C C C C A A A A A G T T A A T C T G T T T G G A C T G	18rs21_al2.seq							
11813	I A A A T C T G C T T R I G A T T T T A I C T G I A A A R G I I A A A I A C C T C C G A G G A A G G	515_al2.seq							
11799	A T A G T C T G T T C A A A G T T T A A T T T C C C C A A A A A G T T A A T C T G T T T G G A C T G	cjb111_al2.seq							
11839	A T A G T C T G T T C A A A G T T T A A T T T C C C C A A A A A G T T A A T C T G T T T G G A C T G	h36b_al2.seq							
A T A T T T C T C T T T C A A A T G T G C T A A T T C A G G T C C G T C T C C T G C A A T C T G T A Majority									
12060		12070		12080		12090		12100	
11951	A T A T T T C T C T T T C A A A T G T G C T A A T T C A G G T C C G T C T C C T G C A A T C T G T A	2603_al2.seq							
11952	A T A T T T C T C T T T C A A A T G T G C T A A T T C A G G T C C G T C T C C T G C A A T C T G T A	18rs21_al2.seq							
11863	G T A G A T A A T A T C C I G A G A A A A A C A G C A A C T G T T I I T A G C T T A T T T T C C A T A	515_al2.seq							
11849	A T A T T T C T C T T T C A A A T G T G C T A A T T C A G G T C C G T C T C C T G C A A T C T G T A	cjb111_al2.seq							
11889	A T A T T T C T C T T T C A A A T G T G C T A A T T C A G G T C C G T C T C C T G C A A T C T G T A	h36b_al2.seq							
A A T A A A C A T T T T C A - G A G T A C T G T G A C A T C G A A A A T G C T T C T A A G A G C A A Majority									
12110		12120		12130		12140		12150	
12001	A A T A A A C A T T T T C A - G A G T A C T G T G A C A T C G A A A A T G C T T C T A A G A G C A A	2603_al2.seq							
12002	A A T A A A C A T T T T C A - G A G T A C T G T G A C A T C G A A A A T G C T T C T A A G A G C A A	18rs21_al2.seq							
11913	T T T A T G C A C T T T C A T G A T A A G C C A T C T T I I A A G C C I T T A A T T A I A G C A A	515_al2.seq							
11899	A A T A A A C A T T T T C A - G A G T A C T G T G A C A T C G A A A A T G C T T C T A A G A G C A A	cjb111_al2.seq							
11939	A A T A A A C A T T T T C A - G A G T A C T G T G A C A T C G A A A A T G C T T C T A A G A G C A A	h36b_al2.seq							

FIGURE 19Z



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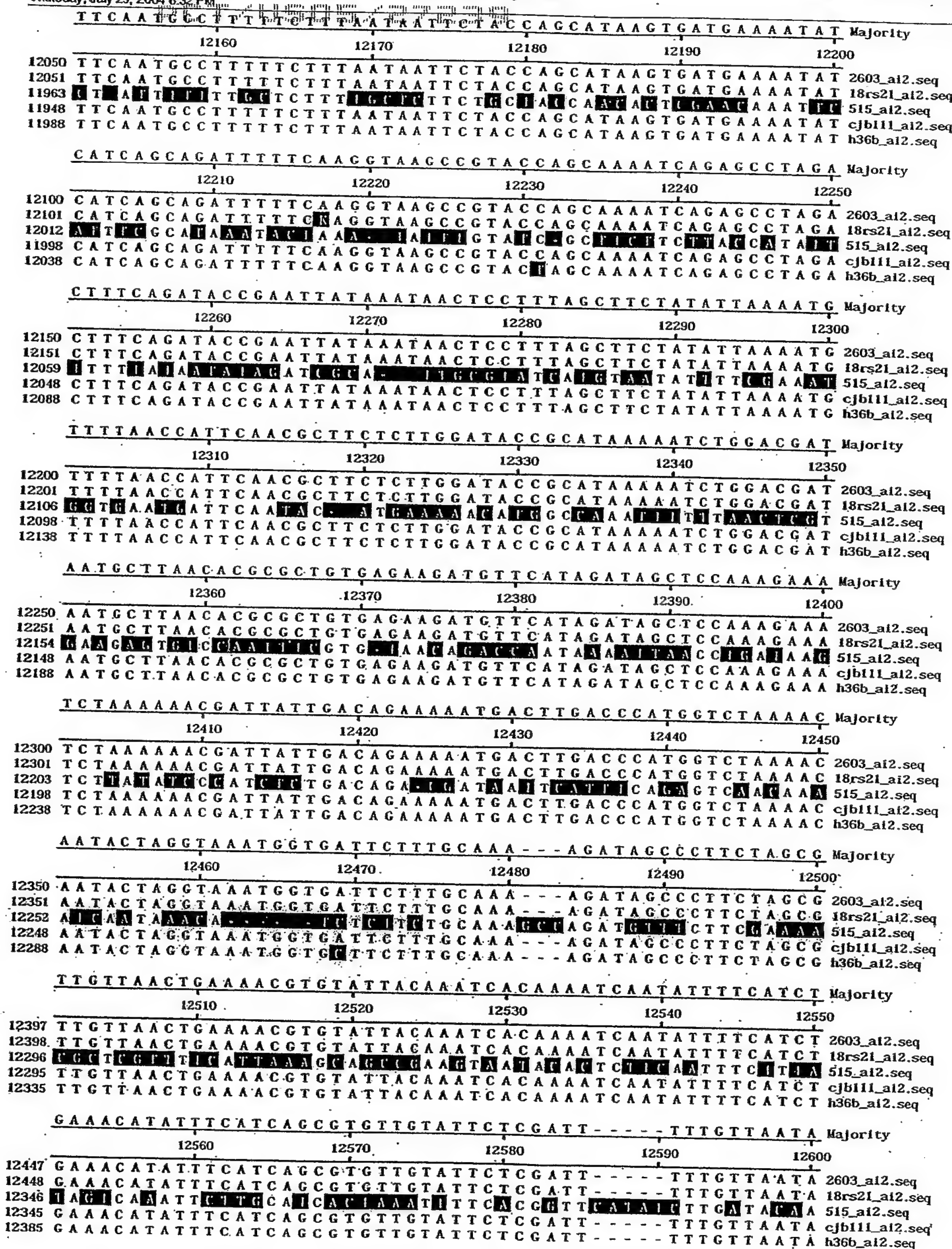


FIGURE 19AA

**FIGURE 19AB.**



**FIGURE 19AC**



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C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G G T G C C C C T C T T C A T T A T A A Majority  
13510 13520 13530 13540 13550

13358 C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G G T G C C C C T C T T C A T T A T A A 2603\_al2.seq  
13359 C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G G T G C C C C T C T T C A T T A T A A 18rs21\_al2.seq  
13270 C T A C C A C T T C G C C A A T G A T T A A C A T T G A G A T A G C A T T G T T A A A A G A A 515\_al2.seq  
13256 C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G G T G C C C C T C T T C A T T A T A A cjb111\_al2.seq  
13296 C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G G T G C C C C T C T T C A T T A T A A h36b\_al2.seq

T C G G C T A C T A A T T G A G A A A T T T C T T C C T T A T T T T T C G A G C C A T T A T C T A C Majority  
13560 13570 13580 13590 13600

13408 T C G G C T A C T A A T T G A G A A A T T T C T T C C T T A T T T T T C G A G C C A T T A T C T A C 2603\_al2.seq  
13409 T C G G C T A C T A A T T G A G A A A T T T C T T C C T T A T T T T T C G A G C C A T T A T C T A C 18rs21\_al2.seq  
13318 C A A G T T T C A A A A G T A A A G C C T T A G C T T G C T T T T C T T T T C C I G A A G A A A A T 515\_al2.seq  
13306 T C G G C C A C T A A T T G A G A A A T T T C T C C T T A T T T T T C G A G C C A T T A T C T A C cjb111\_al2.seq  
13346 T C G G C C A C T A A T T G A G A A A T T T C T T C C T T A T T T T T C G A G C C A T T A T C T A C h36b\_al2.seq

G A T G T A G A T A T G G C T T A C T T G A G G A - - - - T A A A T T G C T C G A A T G T T C T G Majority  
13610 13620 13630 13640 13650

13458 G A T A T A G A T G T G G C T T A C T T G A G G A - - - - T A A A T T G C T C G A A T G T T C T G 2603\_al2.seq  
13459 G A T A T A G A T G T G G C T T A C T T G A G G A - - - - T A A A T T G C T C G A A T G T T C T G 18rs21\_al2.seq  
13368 A G A C T G G A G A G C C T T G G I A A G A A A A C P G C C C G A T T G C T G A A A C A A T G G T 515\_al2.seq  
13356 G A T G T A G A T A T G G C T T A C T T G A G G A - - - - T A A A T T G C T C G A A T G T T C T G cjb111\_al2.seq  
13396 G A T G T A G A T A T G G C T T A C T T G A G G A - - - - T A A A T T G C T C G A A T G T T C T G h36b\_al2.seq

A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T Majority  
13660 13670 13680 13690 13700

13503 A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T 2603\_al2.seq  
13504 A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T 18rs21\_al2.seq  
13418 A A A T A A A A F A C G A A T A A T I I I A T A A A I I G A I C A A A A I A I I C A G G A C P G A 515\_al2.seq  
13401 A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T cjb111\_al2.seq  
13441 A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T h36b\_al2.seq

T C A T G T T C T A T G C T C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C - - - - Majority  
13710 13720 13730 13740 13750

13553 T C A T G T T C T A T G C T C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C - - - - 2603\_al2.seq  
13554 T C A T G T T C T A T G C T C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C - - - - 18rs21\_al2.seq  
13468 G C A A A T A T T G C A I T G C T G C C A G G A T A C T C T T G T T A A G A G A G A I A A A A I C 515\_al2.seq  
13451 T C A T G T T C T A T G C T C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C - - - - cjb111\_al2.seq  
13491 T C A T G T T C T T T G C T C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C - - - - h36b\_al2.seq

- - - - T G G T G C T T T T G G T T A T A A A A A C G A T A C C G A C A T A G A T A G T T A C T G C T Majority  
13760 13770 13780 13790 13800

13598 - - - - T G G T G C T T T T G G T T A T A A A A A C G A T A C C G A C A T A G A T A G T T A C T G C T 2603\_al2.seq  
13599 - - - - T G G T G C T T T T G G T T A T A A A A A C G A T A C C G A C A T A G A T A G T T A C T G C T 18rs21\_al2.seq  
13518 T G C A T G C T G A T T T T G C C G T A A A A A I A A C A I I A A A G T T G G T C I T A G A T G I A 515\_al2.seq  
13496 - - - - T G G T G C T T T T G G T T A T A A A A A C G A T A C C G A C A T A G A T A G T T A C T G C T cjb111\_al2.seq  
13536 - - - - T G G T G C T T T T G G T T A T A A A A A C G A T A C C G A C A T A G A T A G T T A C T G C T h36b\_al2.seq

A C T A A A C T T T G A A T - - - - G A C A T A A T T T A C C A A T G A T A C T G A C A T T T G A G Majority  
13810 13820 13830 13840 13850

13644 A C T A A A C T T T G A A T - - - - G A C A T A A T T T A C C A A T G A T A C T G A C A T T T G A G 2603\_al2.seq  
13645 A C T A A A C T T T G A A T - - - - G A C A T A A T T T A C C A A T G A T A C T G A C A T T T G A G 18rs21\_al2.seq  
13567 I I A A G A T A T T C T A A T T T A C T A A A T G G T A T C I T G A I A A I G A I C A A A T G C A A T T A A A 515\_al2.seq  
13542 A C T A A A C T T T G A A T - - - - G A C A T A A T T T A C C A A T G A T A C T G A C A T T T G A G cjb111\_al2.seq  
13582 A C T A A A C T T T G A A T - - - - G A C A T A A T T T A C C A A T G A T A C T G A C A T T T G A G h36b\_al2.seq

T A T T G A T A T A A T A G A G T A C A G C T C C A C T A A G A - G T A G C A G C A A T T A A A T A Majority  
13860 13870 13880 13890 13900

13690 T A T T G A T A T A A T A G A G T A C A G C T C C A C T A A G A - G T A G C A G C A A T T A A A T A 2603\_al2.seq  
13691 T A T T G A T A T A A T A G A G T A C A G C T C C A C T A A G A - G T A G C A G C A A T T A A A T A 18rs21\_al2.seq  
13617 T G A T G C I A G A C T G I I A A A I I I C G A G I A A I G A I C A A A T G C A A T T A A A 515\_al2.seq  
13588 T A T T G A T A T A A T A G A G T A C A G C T C C A C T A A G A - G T A G C A G C A A T T A A A T A cjb111\_al2.seq  
13628 T A T T G A T A T A A T A A G T A C A G C T C C A C T A A G A - G T A G C A G C A A T T A A A T A h36b\_al2.seq

G C G C A G C A T T C C T C T T G T T A A T T C T T T A A A A G T A A A T A C A T C T C T T A A A G Majority  
13910 13920 13930 13940 13950

13739 G C G C A G C A T T C C T C T T G T T A A T T C T T T A A A A G T A A A T A C A T C T C T T A A A G 2603\_al2.seq  
13740 G C G C A G C A T T C C T C T T G T T A A T T C T T T A A A A G T A A A T A C A T C T C T T A A A G 18rs21\_al2.seq  
13665 - - - - A A C A T A T A A A G T G C T A G G T C A T T T G G C G T C I T A C A A C T A A G A A G 515\_al2.seq  
13637 G C G C A G C A T T C C T C T T G T T A A T T C T T T A A A A G T A A A T A C A T C T C T T A A A G cjb111\_al2.seq  
13677 G C G C A G C A T T C C T C T T G T T A A T T C T T T A A A A G T A A A T A C A T C T C T T A A A G h36b\_al2.seq

FIGURE 19AD





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G T T G C T T C A T A A G T T G T C T G A A G G C C T A A T T C T A A A G T C A C A T G C A T T T C T Majority
 10      20      30      40      50
1  G .....
1  G T T G C T T C A T A A G T T G T C T G A A G G C C T A A T T C T A A A G T C A C A T G C A T T T C T 2603_al2.seq
  nem316_al2.seq
T T C A G A A A G T T C A G C G A G A T A G T A T A T A G T T T C A T C A G G T A A G C A A T C C G Majority
 60      70      80      90     100
2  .....
51 T T C A G A A A G T T C A G C G A G A T A G T A T A T A G T T T C A T C A G G T A A G C A A T C C G 2603_al2.seq
  nem316_al2.seq
G C C T T G T T C C G A T G T T G A T T C C G A T A A C T C C T G G C T C A T T A A T A G C C T G T Majority
 110     120     130     140     150
2  G C C T T G T T C C G A T G T T G A T T C C G A T A A C T C C T G G C T C A T T A A T A G C C T G T 2603_al2.seq
101 G C C T T G T T C C G A T G T T G A T T C C G A T A A C T C C T G G C T C A T T A A T A G C C T G T nem316_al2.seq
T C G T A A C G C T G T T T A A T T A T C T C T A A C T T A G C A T G G G T A T T G G T A A A A T T Majority
 160     170     180     190     200
52 T C G T A A C G C T G T T T A A T T A T C T C T A A C T T A G C A T G G G T A T T G G T A A A A T T 2603_al2.seq
151 T C G T A A C G C T G T T T A A T T A T C T C T A A C T T A G C A T G G G T A T T G G T A A A A T T nem316_al2.seq
T T G A A A A T A G A C T A A G T A T T T A T T A A C C T C G G G C C A C T T T C T A T G C A T G A Majority
 210     220     230     240     250
102 T T G A A A A T A G A C T A A G T A T T T A T T A A C C T C G G G C C A C T T T C T A T G C A T G A 2603_al2.seq
201 T T G A A A A T A G A C T A A G T A T T T A T T A A C C T C G G G C C A C T T T C T A T G C A T G A nem316_al2.seq
A A T C A A T T T C T T T A T A G A A T T G T T C A C G A A T A G G A G C T T C T G G A G C A A C T Majority
 260     270     280     290     300
152 A A T C A A T T T C T T T A T A G A A T T G T T C A C G A A T A G G A G C T T C T G G A G C A A C T 2603_al2.seq
251 A A T C A A T T T C T T T A T A G A A T T G T T C A C G A A T A G G A G C T T C T G G A G C A A C T nem316_al2.seq
A T A G C A T C C C C T G A A C C A G A A A C T G T G C A A A A A G T G C A T C C T C C T C T A G C Majority
 310     320     330     340     350
202 A T A G C A T C C C C T G A A C C A G A A A C T G T G C A A A A A G T G C A T C C T C C T C T A G C 2603_al2.seq
301 A T A G C A T C C C C T G A A C C A G A A A C T G T G C A A A A A G T G C A T C C T C C T C T A G C nem316_al2.seq
A A C T G T T C C G T C T C T G T T A G G A C A G T C A A A A C C A G C A T C T A T A G G T A A T T Majority
 360     370     380     390     400
252 A A C T G T T C C G T C T C T G T T A G G A C A G T C A A A A C C A G C A T C T A T A G G T A A T T 2603_al2.seq
351 A A C T G T T C C G T C T C T G T T A G G A C A G T C A A A A C C A G C A T C T A T A G G T A A T T nem316_al2.seq
T A A A T A T T T T T T C T C C A A A G A G T T C T C G A T A A T A A T C A T T A A T C G C A C G A Majority
 410     420     430     440     450
302 T A A A T A T T T T T T C T C C A A A G A G T T C T C G A T A A T A A T C A T T A A T C G C A C G A 2603_al2.seq
401 T A A A T A T T T T T T C T C C A A A G A G T T C T C G A T A A T A A T C A T T A A T C G C A C G A nem316_al2.seq
T A A C G T T T T T T C A T A G G A T A A T T G T A T C A C A A T T T T A A C T A A A A T A A C C T Majority
 460     470     480     490     500
352 T A A C G T T T T T T C A T A G G A T A A T T G T A T C A C A A T T T T A A C T A A A A T A A C C T 2603_al2.seq
451 T A A C G T T T T T T C A T A G G A T A A T T G T A T C A C A A T T T T A A C T A A A A T A A C C T nem316_al2.seq
C A C T A C T A C A A T A A A A C T T A A A A A G A T T G G A A C G T C A G T T A G T C C C A A T C Majority
 510     520     530     540     550
402 C A C T A C T A C A A T A A A A C T T A A A A A G A T T G G A A C G T C A G T T A G T C C C A A T C 2603_al2.seq
501 C A C T A C T A C A A T A A A A C T T A A A A A G A T T G G A A C G T C A G T T A G T C C C A A T C nem316_al2.seq
T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C G Majority
 560     570     580     590     600
452 T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C G 2603_al2.seq
551 T T T T A T T T A C T T C A C T T T C T T T A A C C A A T C C T T G G C T A A A A A G A T A T A C G nem316_al2.seq
C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C T Majority
 610     620     630     640     650
502 C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C T 2603_al2.seq
601 C A G T T A G A T T C A A A A T A C C A T A A G C A A G T A T A A A A C C A G C T A A A A C A T C T nem316_al2.seq
```

Figure 20



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	660	670	680	690	700	Majority
552	GTCGGGAAAATGAACCCCTAGGTAATAACGAGATAACCCAATTAAAAAAAT					2603_al2.seq
651	GTCGGGAAAATGAACCCCTAGGTAATAACGAGATAACCCAATTAAAAAAAT					nem316_al2.seq
	710	720	730	740	750	Majority
602	GAGCAAACCCAAAGTACCTTGGGCACAACAGTTTCCATATACTCTTAGGCCA					2603_al2.seq
701	GAGCAAACCCAAAGTACCTTGGGCACAACAGTTTCCATATACTCTTAGGCCA					nem316_al2.seq
	760	770	780	790	800	Majority
652	TATAGTACTGCAATAAAATAATAATACTCCCAAATATCATAAATGTTCCC					2603_al2.seq
751	TATAGTACTGCAATAAAATAATAATACTCCCAAATATCATAAATGTTCCC					nem316_al2.seq
	810	820	830	840	850	Majority
702	ATCGAGTGCCCACTGGGGAACGAATAGCCACCTGCAAATACTAAATGGGT					2603_al2.seq
801	ATCGAGTGCCCACTGGGGAACGAATAGCCACCTGCAAATACTAAATGGGT					nem316_al2.seq
	860	870	880	890	900	Majority
752	TAAAGTTGGTCTTACTCTTTGAAAAATAAGTTTTAAAGAAAGTATACATA					2603_al2.seq
851	TAAAGTTGGTCTTACTCTTTGAAAAATAAGTTTTAAAGAAAGTATACATA					nem316_al2.seq
	910	920	930	940	950	Majority
802	TACCAGAGATAAATAGCATTCTACTGCGATAAATCTAGCTTGAGGATACCAC					2603_al2.seq
901	TACCAGAGATAAATAGCATTCTACTGCGATAAATCTAGCTTGAGGATACCAC					nem316_al2.seq
	960	970	980	990	1000	Majority
852	TTCTTAAGGTAACAGAAAGTGACGCTCATAATCGCAATAGCTATCTGGCT					2603_al2.seq
951	TTCTTAAGGTAACAGAAAGTGACGCTCATAATCGCAATAGCTATCTGGCT					nem316_al2.seq
	1010	1020	1030	1040	1050	Majority
902	TACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGAT					2603_al2.seq
1001	TACAGTATTACCAATCACAGTGATTAACTTGAAAAATCTTGTAGAAAGAT					nem316_al2.seq
	1060	1070	1080	1090	1100	Majority
952	TTGGCAACTGTCCTCTAACACTTTCTTGAATGTTTTGGTCAAATGCAATT					2603_al2.seq
1051	TTGGCAACTGTCCTCTAACACTTTCTTGAATGTTTTGGTCAAATGCAATT					nem316_al2.seq
	1110	1120	1130	1140	1150	Majority
1002	ACAGTGTGCGGGCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGAT					2603_al2.seq
1101	ACAGTGTGCGGGCCAATATTTGATGACCAATCCTAAACTGAAAAATAAGAT					nem316_al2.seq
	1160	1170	1180	1190	1200	Majority
1052	AATAGCAATAAATGCTTGAATAAGTTTACTATTTTGACGAGATAACATTA					2603_al2.seq
1151	AATAGCAATAAATGCTTGAATAAGTTTACTATTTTGACGAGATAACATTA					nem316_al2.seq
	1210	1220	1230	1240	1250	Majority
1102	GTCTTTTTTATATCTTTCTAATAATTGGCAAACAAGCCACGTAAGTTAGATA					2603_al2.seq
1201	GTCTTTTTTATATCTTTCTAATAATTGGCAAACAAGCCACGTAAGTTAGATA					nem316_al2.seq
	1260	1270	1280	1290	1300	Majority
1152	GAAAAACAATCGAAATTAAAAATTCCCTCAACGATATTAAATGGAATAACCA					2603_al2.seq
1251	GAAAAACAATCGAAATTAAAAATTCCCTCAACGATATTAAATGGAATAACCA					nem316_al2.seq

FIGURE 20A

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	T T G T T A A A A G G T A A T T G C C T A C A C C A A T A A A T G T T C T G A T A T C A A A G T T A	Majority
	1310 1320 1330 1340 1350	
1202	T T G T T A A A A G G T A A T T G C C T A C A C C A A T A A A T G T T C T G A T A T C A A A G T T A	2603_al2.seq
1301	T T G T T A A A A G G T A A T T G C C T A C A C C A A T A A A T G T T C T G A T A T C A A A G T T A	nem316_al2.seq
	G C A A A T A T A G C A T A C A A A G G A A T C G C A A A G A C A T A G T T G A G A G C T A C C A T	Majority
	1360 1370 1380 1390 1400	
1252	G C A A A T A T A G C A T A C A A A G G A A T C G C A A A G A C A T A G T T G A G A G C T A C C A T	2603_al2.seq
1351	G C A A A T A T A G C A T A C A A A G G A A T C G C A A A G A C A T A G T T G A G A G C T A C C A T	nem316_al2.seq
	A G A T A C G G T C A A G C T A A C T G T A C C A A A T A G A C T A G C T T T A A T A A A A T C T T	Majority
	1410 1420 1430 1440 1450	
1302	A G A T A C G G T C A A G C T A A C T G T A C C A A A T A G A C T A G C T T T A A T A A A A T C T T	2603_al2.seq
1401	A G A T A C G G T C A A G C T A A C T G T A C C A A A T A G A C T A G C T T T A A T A A A A T C T T	nem316_al2.seq
	T T G C A C T C T C T C T A T T T T T C C A G A A A A T A G C G A A A C T T G C T A A A A A T A A A	Majority
	1460 1470 1480 1490 1500	
1352	T T G C A C T C T C T C T A T T T T T C C A G A A A A T A G C G A A A C T T G C T A A A A A T A A A	2603_al2.seq
1451	T T G C A C T C T C T C T A T T T T T C C A G A A A A T A G C G A A A C T T G C T A A A A A T A A A	nem316_al2.seq
	G C T A G A G C A A C C A T A T T C A T C G G T A A A C C G A T A A A G G T T T C T G G A C C A C G	Majority
	1510 1520 1530 1540 1550	
1402	G C T A G A G C A A C C A T A T T C A T C G G T A A A C C G A T A A A G G T T T C T G G A C C A C G	2603_al2.seq
1501	G C T A G A G C A A C C A T A T T C A T C G G T A A A C C G A T A A A G G T T T C T G G A C C A C G	nem316_al2.seq
	A T T A G C A A G T A T A A C T T T T A A A A G T G A T C T T A A T A A G A G T A C A C C A T A A C	Majority
	1560 1570 1580 1590 1600	
1452	A T T A G C A A G T A T A A C T T T T A A A A G T G A T C T T A A T A A G A G T A C A C C A T A A C	2603_al2.seq
1551	A T T A G C A A G T A T A A C T T T T A A A A G T G A T C T T A A T A A G A G T A C A C C A T A A C	nem316_al2.seq
	T T G A T T T C A A A T C A A A T A A A A T A A A A G C A A C T A A C A T C G G A A G G A T T G A A	Majority
	1610 1620 1630 1640 1650	
1502	T T G A T T T C A A A T C A A A T A A A A T A A A A G C A A C T A A C A T C G G A A G G A T T G A A	2603_al2.seq
1601	T T G A T T T C A A A T C A A A T A A A A T A A A A G C A A C T A A C A T C G G A A G G A T T G A A	nem316_al2.seq
	A A A T C A A C C T T T A A A A A T T C T G C T C C T G G T A T T A A T G G A A A T G A A A C C A T	Majority
	1660 1670 1680 1690 1700	
1552	A A A T C A A C C T T T A A A A A T T C T G C T C C T G G T A T T A A T G G A A A T G A A A C C A T	2603_al2.seq
1651	A A A T C A A C C T T T A A A A A T T C T G C T C C T G G T A T T A A T G G A A A T G A A A C C A T	nem316_al2.seq
	C A T C A A T A C A A A A G A T A A G G C A G A A A G A A T G G C G A T T G T C A C C A T T T T A C	Majority
	1710 1720 1730 1740 1750	
1602	C A T C A A T A C A A A A G A T A A G G C A G A A A G A A T G G C G A T T G T C A C C A T T T T A C	2603_al2.seq
1701	C A T C A A T A C A A A A G A T A A G G C A G A A A G A A T G G C G A T T G T C A C C A T T T T A C	nem316_al2.seq
	G T G T A T T T G T C A T A A A A A A A T T C C T C C A A T T T A A A T A A A T T G A A A G A A G C	Majority
	1760 1770 1780 1790 1800	
1652	G T G T A T T T G T C A T A A A A A A A T T C C T C C A A T T T A A A T A A A T T G A A A G A A G C	2603_al2.seq
1751	G T G T A T T T G T C A T A A A A A A A T T C C T C C A A T T T A A A T A A A T T G A A A G A A G C	nem316_al2.seq
	T C C A A A G G T A A G C G T A T G T A C G C G A A A A A A C C T T T G T C T T C T C C C A T C C	Majority
	1810 1820 1830 1840 1850	
1702	T C C A A A G G T A A G C G T A T G T A C G C G A A A A A A C C T T T G T C T T C T C C C A T C C	2603_al2.seq
1801	T C C A A A G G T A A G C G T A T G T A C G C G A A A A A A A C C T T T G T C T T C T C C C A T C C	nem316_al2.seq
	A G A C T T T A C T G T C G G T T G T G G A A T C T C A C C A C A T C A G C T T T C G C T C G C G G	Majority
	1860 1870 1880 1890 1900	
1751	A G A C T T T A C T G T C G G T T G T G G A A T C T C A C C A C A T C A G C T T T C G C T C G C G G	2603_al2.seq
1851	A G A C T T T A C T G T C G G T T G T G G A A T C T C A C C A C A T C A G C T T T C G C T C G C G G	nem316_al2.seq
	A C T G A T G C T T C A C A A C T G A C A A A T A A G T T G G A A G C G A T T A C C G C C G G T C G	Majority
	1910 1920 1930 1940 1950	
1801	A C T G A T G C T T C A C A A C T G A C A A A T A A G T T G G A A G C G A T T A C C G C C G G T C G	2603_al2.seq
1901	A C T G A T G C T T C A C A A C T G A C A A A T A A G T T G G A A G C G A T T A C C G C C G G T C G	nem316_al2.seq

FIGURE 20B



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	GGAATTACACCCCTGCCCTGAAGACACCTATAGCATAACAAAAAAACTTTG	Majority
	1960 1970 1980 1990 2000	
1851	GGAATTACACCCCTGCCCTGAAGACACCTATAGCATAACAAAAAAACTTTG	2603_al2.seq
1951	GGAATTACACCCCTGCCCTGAAGACACCTATAGCATAACAAAAAAACTTTG	nem316_al2.seq
	CAATTGCAAGTTTTTTTAATTACTAATTAGTAGTAGTGATTAAAAATCATA	Majority
	2010 2020 2030 2040 2050	
1901	CAATTGCAAGTTTTTTTAATTACTAATTAGTAGTAGTGATTAAAAATCATA	2603_al2.seq
2001	CAATTGCAAGTTTTTTTAATTACTAATTAGTAGTAGTGATTAAAAATCATA	nem316_al2.seq
	TTAATACCAAATTACTATGCTGTATCGTTTCTTTTCAGATTTGCTATTTTTT	Majority
	2060 2070 2080 2090 2100	
1951	TTAATACCAAATTACTATGCTGTATCGTTTCTTTTCAGATTTGCTATTTTTT	2603_al2.seq
2051	TTAATACCAAATTACTATGCTGTATCGTTTCTTTTCAGATTTGCTATTTTTT	nem316_al2.seq
	AGTTTTTCTTAAAAAGATAAAACAAAATTCCCAAATAATACAACCAAGAA	Majority
	2110 2120 2130 2140 2150	
2001	AGTTTTTCTTAAAAAGATAAAACAAAATTCCCAAATAATACAACCAAGAA	2603_al2.seq
2101	AGTTTTTCTTAAAAAGATAAAACAAAATTCCCAAATAATACAACCAAGAA	nem316_al2.seq
	TTGTCAGTCCTCCACCAATAATCATTCTGTTTTAGGAAGAAATGATTGT	Majority
	2160 2170 2180 2190 2200	
2051	TTGTCAGTCCTCCACCAATAATCATTCTGTTTTAGGAAGAAATGATTGT	2603_al2.seq
2151	TTGTCAGTCCTCCACCAATAATCATTCTGTTTTAGGAAGAAATGATTGT	nem316_al2.seq
	GGAAAAAGCGGTTGTGATGGTTTAGGATTTGTTGGTGGAGGAGTTTCTTT	Majority
	2210 2220 2230 2240 2250	
2101	GGAAAAAGCGGTTGTGATGGTTTAGGATTTGTTGGTGGAGGAGTTTCTTT	2603_al2.seq
2201	GGAAAAAGCGGTTGTGATGGTTTAGGATTTGTTGGTGGAGGAGTTTCTTT	nem316_al2.seq
	TTCTGTTTTCTACCTCTACTTCTCTGTTTTATTAGCAACTACAGCAACTA	Majority
	2260 2270 2280 2290 2300	
2151	TTCTGTTTTCTACCTCTACTTCTCTGTTTTATTAGCAACTACAGCAACTA	2603_al2.seq
2251	TTCTGTTTTCTACCTCTACTTCTCTGTTTTATTAGCAACTACAGCAACTA	nem316_al2.seq
	CAGCATCCTTCATAGATATACGGTAACCAGTTAGTGCTTTTGTCTTCTCGA	Majority
	2310 2320 2330 2340 2350	
2201	CAGCATCCTTCATAGATATACGGTAACCAGTTAGTGCTTTTGTCTTCTCGA	2603_al2.seq
2301	CAGCATCCTTCATAGATATACGGTAACCAGTTAGTGCTTTTGTCTTCTCGA	nem316_al2.seq
	AAAATATACTTACCAGGTAATAAACCTTCAACCTCAATTTCTCCCTTATC	Majority
	2360 2370 2380 2390 2400	
2251	AAAATATACTTACCAGGTAATAAACCTTCAACCTCAATTTCTCCCTTATC	2603_al2.seq
2351	AAAATATACTTACCAGGTAATAAACCTTCAACCTCAATTTCTCCCTTATC	nem316_al2.seq
	ATCAGTTACTAATGAAGTAATCCCATCTTGATCGGTCGTAAATCGTCCAT	Majority
	2410 2420 2430 2440 2450	
2301	ATCAGTTACTAATGAAGTAATCCCATCTTGATCGGTCGTAAATCGTCCAT	2603_al2.seq
2401	ATCAGTTACTAATGAAGTAATCCCATCTTGATCGGTCGTAAATCGTCCAT	nem316_al2.seq
	TTTTAAAGCGAACTGGCTGATTCTGGTTATCGTATAATAACAATATTACT	Majority
	2460 2470 2480 2490 2500	
2351	TTTTAAAGCGAACTGGCTGATTCTGGTTATCGTATAATAACAATATTACT	2603_al2.seq
2451	TTTTAAAGCGAACTGGCTGATTCTGGTTATCGTATAATAACAATATTACT	nem316_al2.seq
	CCTGATAGCCTTTTCTTTATCTTTCCTTCTTTTGTATATTTAATAAGTTT	Majority
	2510 2520 2530 2540 2550	
2401	CCTGATAGCCTTTTCTTTATCTTTCCTTCTTTTGTATATTTAATAAGTTT	2603_al2.seq
2501	CCTGATAGCCTTTTCTTTATCTTTCCTTCTTTTGTATATTTAATAAGTTT	nem316_al2.seq
	TAAATCGGCCCTGTTTCAACTTTTTCGCTTAGGATTTATCTGTAATTGATTTC	Majority
	2560 2570 2580 2590 2600	
2451	TAAATCGGCCCTGTTTCAACTTTTTCGCTTAGGATTTATCTGTAATTGATTTC	2603_al2.seq
2551	TAAATCGGCCCTGTTTCAACTTTTTCGCTTAGGATTTATCTGTAATTGATTTC	nem316_al2.seq

FIGURE 20C

		ATAACTTATCATCTGGTATTTCAATATAAAAAAGGTACTATTGTTGAAACG	Majority
		2610 2620 2630 2640 2650	
2501		ATAACTTATCATCTGGTATTTCAATATAAAAAAGGTACTATTGTTGAAACG	2603_al2.seq
2601		ATAACTTATCATCTGGTATTTCAATATAAAAAAGGTACTATTGTTGAAACG	nem316_al2.seq
		CTTTGATCAGCTTTTATAAGCACGACCAAAAGTACGAACCATTTGGGAGTGC	Majority
		2660 2670 2680 2690 2700	
2551		CTTTGATCAGCTTTTATAAGCACGACCAAAAGTACGAACCATTTGGGAGTGC	2603_al2.seq
2651		CTTTGATCAGCTTTTATAAGCACGACCAAAAGTACGAACCATTTGGGAGTGC	nem316_al2.seq
		TATCTTTGTCTGACCATTAGTATCAGTAGGAGAAGTCAAGATACTCTTAT	Majority
		2710 2720 2730 2740 2750	
2601		TATCTTTGTCTGACCATTAGTATCAGTAGGAGAAGTCAAGATACTCTTAT	2603_al2.seq
2701		TATCTTTGTCTGACCATTAGTATCAGTAGGAGAAGTCAAGATACTCTTAT	nem316_al2.seq
		ACTTCTGGTTCAATTTCGCTATCTGTCAATTTGGCTCAATAAATCAACTTTT	Majority
		2760 2770 2780 2790 2800	
2651		ACTTCTGGTTCAATTTCGCTATCTGTCAATTTGGCTCAATAAATCAACTTTT	2603_al2.seq
2751		ACTTCTGGTTCAATTTCGCTATCTGTCAATTTGGCTCAATAAATCAACTTTT	nem316_al2.seq
		AAGTTGTCAGTCACAGTCCATAAACGATAAGAAATCCCCTCCTCTGTAGT	Majority
		2810 2820 2830 2840 2850	
2701		AAGTTGTCAGTCACAGTCCATAAACGATAAGAAATCCCCTCCTCTGTAGT	2603_al2.seq
2801		AAGTTGTCAGTCACAGTCCATAAACGATAAGAAATCCCCTCCTCTGTAGT	nem316_al2.seq
		ATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCAG	Majority
		2860 2870 2880 2890 2900	
2751		ATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCAG	2603_al2.seq
2851		ATTTGGCTGAAGTCCTATCTGTGTGATTGTTAGTTGATTAGGGGTATCAG	nem316_al2.seq
		CATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAAG	Majority
		2910 2920 2930 2940 2950	
2801		CATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAAG	2603_al2.seq
2901		CATTTACACTGGCTACCGAAAAAAACGCTAATTGTACCAATCCTAAAAAG	nem316_al2.seq
		CAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATTT	Majority
		2960 2970 2980 2990 3000	
2851		CAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATTT	2603_al2.seq
2951		CAACATAGTAGAAGTCCTAAACTTTTTCTAATCTTTTTTCATTTTTTGATTT	nem316_al2.seq
		CCCTTTCTTTTTCTCTCTTTTAAATTTTCGTTTTTAAATATAATAGTAAAGC	Majority
		3010 3020 3030 3040 3050	
2901		CCCTTTCTTTTTCTCTCTTTTAAATTTTCGTTTTTAAATATAATAGTAAAGC	2603_al2.seq
3001		CCCTTTCTTTTTCTCTCTTTTAAATTTTCGTTTTTAAATATAATAGTAAAGC	nem316_al2.seq
		GACTAATATAAGAATAAAGTATGATAAGAGGAAATAAAGTTTATAGT	Majority
		3060 3070 3080 3090 3100	
2951		GACTAATATAAGAATAAAGTATGATAAGAGGAAATAAAGTTTATAGT	2603_al2.seq
3051		GACTAATATAAGAATAAAGTATGATAAGAGGAAATAAAGTTTATAGT	nem316_al2.seq
		GTGTTTGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTACA	Majority
		3110 3120 3130 3140 3150	
3001		GTGTTTGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTACA	2603_al2.seq
3101		GTGTTTGCAATTCTTTTCATTAAATAGTTCTTTTCTTTTAAACAGGAGGTACA	nem316_al2.seq
		TACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAATA	Majority
		3160 3170 3180 3190 3200	
3051		TACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAATA	2603_al2.seq
3151		TACTTGATTTCGATGCCCTCTAACTAGTAAACGATGTGAATTAATCGAATA	nem316_al2.seq
		AGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAATA	Majority
		3210 3220 3230 3240 3250	
3101		AGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAATA	2603_al2.seq
3201		AGGTGTACATGTTAGCAAAGTCCGATAATCCTTACCTTTTAAACAACCAATA	nem316_al2.seq

FIGURE 20D



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      A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A G Majority
      3260          3270          3280          3290          3300
3151 A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A G 2603_al2.seq
3251 A T T T A G A A A A A T T A T C T G G C T T T A C A A C A C T T A T T T G A T C A A C C T T A T A G nem316_al2.seq

      G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A G Majority
      3310          3320          3330          3340          3350
3201 G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A G 2603_al2.seq
3301 G C T A A A A C T T C T T T G A T A T T A T G A A T A T A A A A A A T T T T T C C T T T T T T A A G nem316_al2.seq

      T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T G Majority
      3360          3370          3380          3390          3400
3251 T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T G 2603_al2.seq
3351 T T T A T C T A A A T C T G T A A A T A A C T T A G C T T T A G G T A A G C C G C G A T G A G C T G nem316_al2.seq

      T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C A Majority
      3410          3420          3430          3440          3450
3301 T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G G A G G T T C C T T C A 2603_al2.seq
3401 T G A T A A C A G T A T G T G A A C T T T T T C C A C C A A T T G G C A A G A G A G T T C C T T C A nem316_al2.seq

      A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A T Majority
      3460          3470          3480          3490          3500
3351 A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A T 2603_al2.seq
3451 A G G T G T C C T G C T C C T T T T T C A A G A A C A C T A C T G G T A G T C C C C G C A T A G A T nem316_al2.seq

      A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T C A G Majority
      3510          3520          3530          3540          3550
3401 A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T C A G 2603_al2.seq
3501 A G G T A A T T T T T G C T T G A T A G A C G G T A T A T C A A T A T A T C C A A T C A T T T C A G nem316_al2.seq

      T A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C A Majority
      3560          3570          3580          3590          3600
3451 T A A T C T C A A G C A T G T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C A 2603_al2.seq
3551 T A A T C T C A A G C A T T G G G C G T A T T C A G C A A T A C C T T T T T T T C T T T T T C A nem316_al2.seq

      G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T T Majority
      3610          3620          3630          3640          3650
3501 G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T T 2603_al2.seq
3601 G T A T A G G G A T C T G A T A G G C G G C T T G G G T C C A G T G T T C T A T T A T A A G C T T T nem316_al2.seq

      T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T T Majority
      3660          3670          3680          3690          3700
3551 T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T T 2603_al2.seq
3651 T G C T A A C T C A A A T C G T C T A T T A A T C T C T T T A G T A T T T A A T T T T T G G G T T T nem316_al2.seq

      G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A A Majority
      3710          3720          3730          3740          3750
3601 G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A A 2603_al2.seq
3701 G A T T A T C A A A G T T A G T T A C T T G A T T A T T A G C T T T A A T A T T A T A G T A C C A A nem316_al2.seq

      T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A G Majority
      3760          3770          3780          3790          3800
3651 T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A G 2603_al2.seq
3751 T T T G A A A T A A A A G G A T A T G A G G T T A T C A A A A G A C C A A C T A A G A A C A A T A G nem316_al2.seq

      T A T C A G G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G T Majority
      3810          3820          3830          3840          3850
3701 T A T C A G G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G T 2603_al2.seq
3801 T A T C A G G C C T A C A T T C A T C C A T C G A T T T A A A A C G A C C G A T T T C T T A A G G T nem316_al2.seq

      T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T G Majority
      3860          3870          3880          3890          3900
3751 T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T G 2603_al2.seq
3851 T T T T C T G A A A T T T T C C T C C C A T T A T G A T T C A A T T C C T T T T C T A A C A C T T G nem316_al2.seq

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FIGURE 20E

FIGURE 20F



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	ATACGTCGATTGATTTCCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAAA	Majority
	4560 4570 4580 4590 4600	
4451	ATACGTCGATTGATTTCCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAAA	2603_al2.seq
4551	ATACGTCGATTGATTTCCTTTCTGACTAAGTTTTTTTAGCAGCTCTCTCAAA	nem316_al2.seq
	ATCCTGTGTTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACCA	Majority
	4610 4620 4630 4640 4650	
4501	ATCCTGTGTTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACCA	2603_al2.seq
4601	ATCCTGTGTTTGATTATTAGATTCTATCGTATAGTAAAAACGTGATACCA	nem316_al2.seq
	CTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGGA	Majority
	4660 4670 4680 4690 4700	
4551	CTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGGA	2603_al2.seq
4651	CTGGATACAATAAAATAGATAGACCTATTAGAAAAAGAATGATAAAAGGA	nem316_al2.seq
	AGATTTGACTTCTTCTTTTTTTTTTGTGTTTGTGATTGTTTCTT	Majority
	4710 4720 4730 4740 4750	
4601	AGATTTGACTTCTTCTTTTTTTTTTGTGTTTGTGATTGTTTCTT	2603_al2.seq
4701	AGATTTGACTTCTTCTTTTTTTTGTGTTTGTGATTGTTTCTT	nem316_al2.seq
	CACGTCATCTCCTAGATAATGGCTCTTGCTTATGATCTAAGAGTACTTCT	Majority
	4760 4770 4780 4790 4800	
4651	CACGTCATCTCCTAGATAATGGCTCTTGCTTATGATCTAAGAGTACTTCT	2603_al2.seq
4748	CACGTCATCTCCTAATAATGGCTCTTGCTTATGATCTAAGAGTACTTCT	nem316_al2.seq
	ACTGAAATACCCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACATC	Majority
	4810 4820 4830 4840 4850	
4701	ACTGAAATACCCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACATC	2603_al2.seq
4798	ACTGAAATACCCCTTAGATCATAAGCACAGCTTTAACTGTGCTTATACATC	nem316_al2.seq
	ATCAAAGACTAGCCCTTAAGCTTCTTTGATTGGCGTTTTCATGATAAC	Majority
	4860 4870 4880 4890 4900	
4751	ATCAAAGACTAGCCCTTAAGCTTCTTTGATTGGCGTTTTCATGATAAC	2603_al2.seq
4848	ATCAAAGACTAGCCCTTAAGCTTCTTTGATTGGCGTTTTCATGATAAC	nem316_al2.seq
	TACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTAC	Majority
	4910 4920 4930 4940 4950	
4801	TACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTAC	2603_al2.seq
4898	TACTGCTCCAAGCATAATGCTTAAACCAATAATTGTGAAAAGAATTGTAC	nem316_al2.seq
	CAATACCACCTGTTTGTGGGATTGTTACTTTTTTGTGTTTGTACTTGTG	Majority
	4960 4970 4980 4990 5000	
4851	CAATACCACCTGTTTGTGGGATTGTTACTTTTTTGTGTTTGTACTTGTG	2603_al2.seq
4948	CAATACCACCTGTTTGTGGGATTGTTACTTTTTTGTGTTTGTACTTGTG	nem316_al2.seq
	GCATCTTTTTTTTACAGGTTTTTTTGTGTTATCTGCGTTGTCAGTTTATAGCCCC	Majority
	5010 5020 5030 5040 5050	
4901	GCATCTTTTTTTTACAGGTTTTTTTGTGTTATCTGCGTTGTCAGTTTATAGCCCC	2603_al2.seq
4998	GCATCTTTTTTTTACAGGTTTTTTTGTGTTATCTGCGTTGTCAGTTTATAGCCCC	nem316_al2.seq
	TTTTCTGTATGATGTTTGAATTTACTTCAAAGTTTATATTACCTGCCAATT	Majority
	5060 5070 5080 5090 5100	
4945	TTTTCTGTATGATGTTTGAATTTACTTCAAAGTTTATATTACCTGCCAATT	2603_al2.seq
5048	TTTTCTGTATGATGTTTGAATTTACTTCAAAGTTTATATTACCTGCCAATT	nem316_al2.seq
	TCCGATATCCTGCTGCTGCTTGTGTTTCTTCCAGGTTGTAAGTGCCTTTT	Majority
	5110 5120 5130 5140 5150	
4995	TCCGATATCCTGCTGCTGCTTGTGTTTCTTCCAGGTTGTAAGTGCCTTTT	2603_al2.seq
5098	TCCGATATCCTGCTGCTGCTTGTGTTTCTTCCAGGTTGTAAGTGCCTTTT	nem316_al2.seq
	TCCAGACCTGTAATTTCAAATTGACCTTGGTCTTTGAGGTGTAATTTAAT	Majority
	5160 5170 5180 5190 5200	
5045	TCCAGACCTGTAATTTCAAATTGACCTTGGTCTTTGAGGTGTAATTTAAT	2603_al2.seq
5148	TCCAGACCTGTAATTTCAAATTGACCTTGGTCTTTGAGGTGTAATTTAAT	nem316_al2.seq

FIGURE 20G

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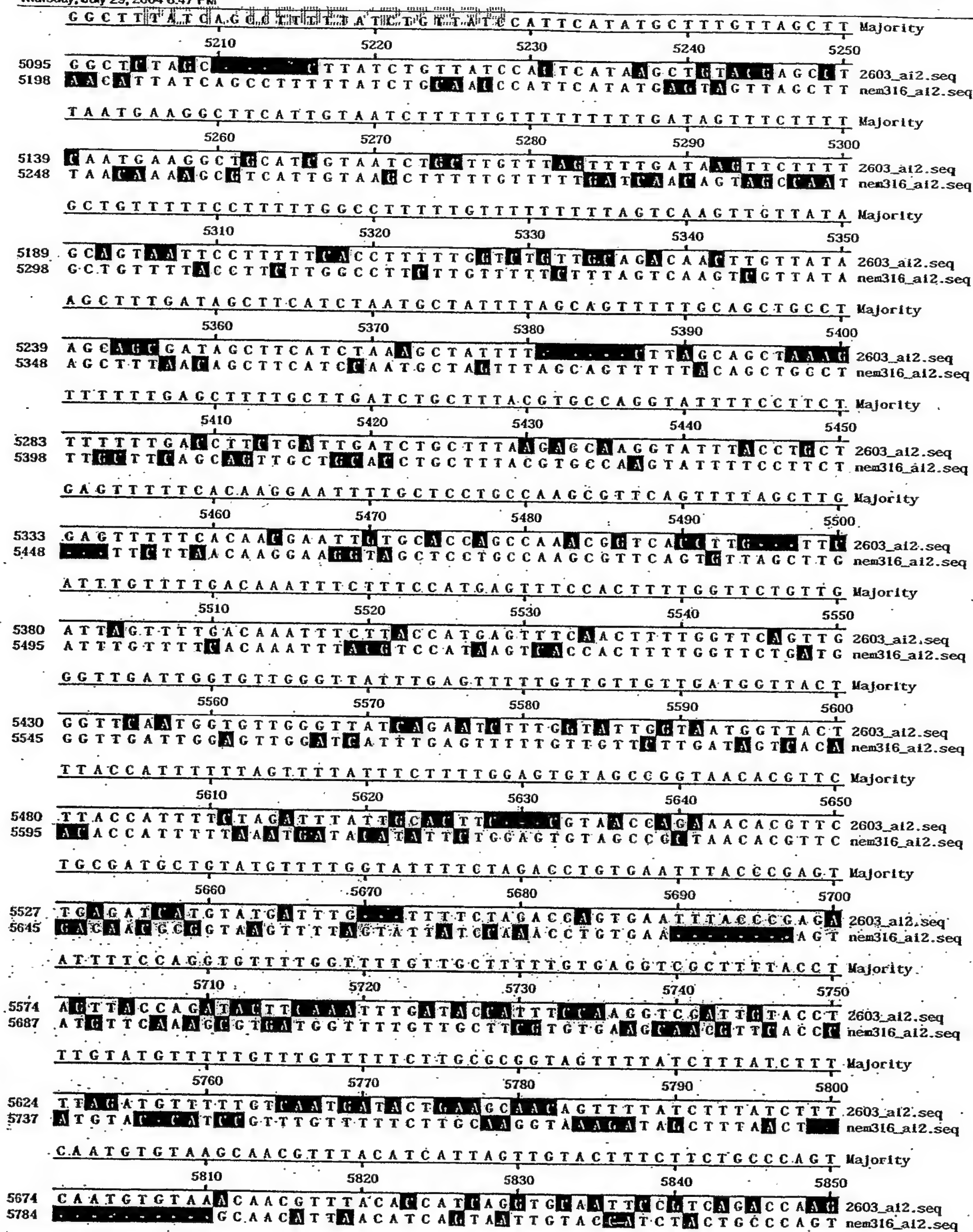


FIGURE 20H



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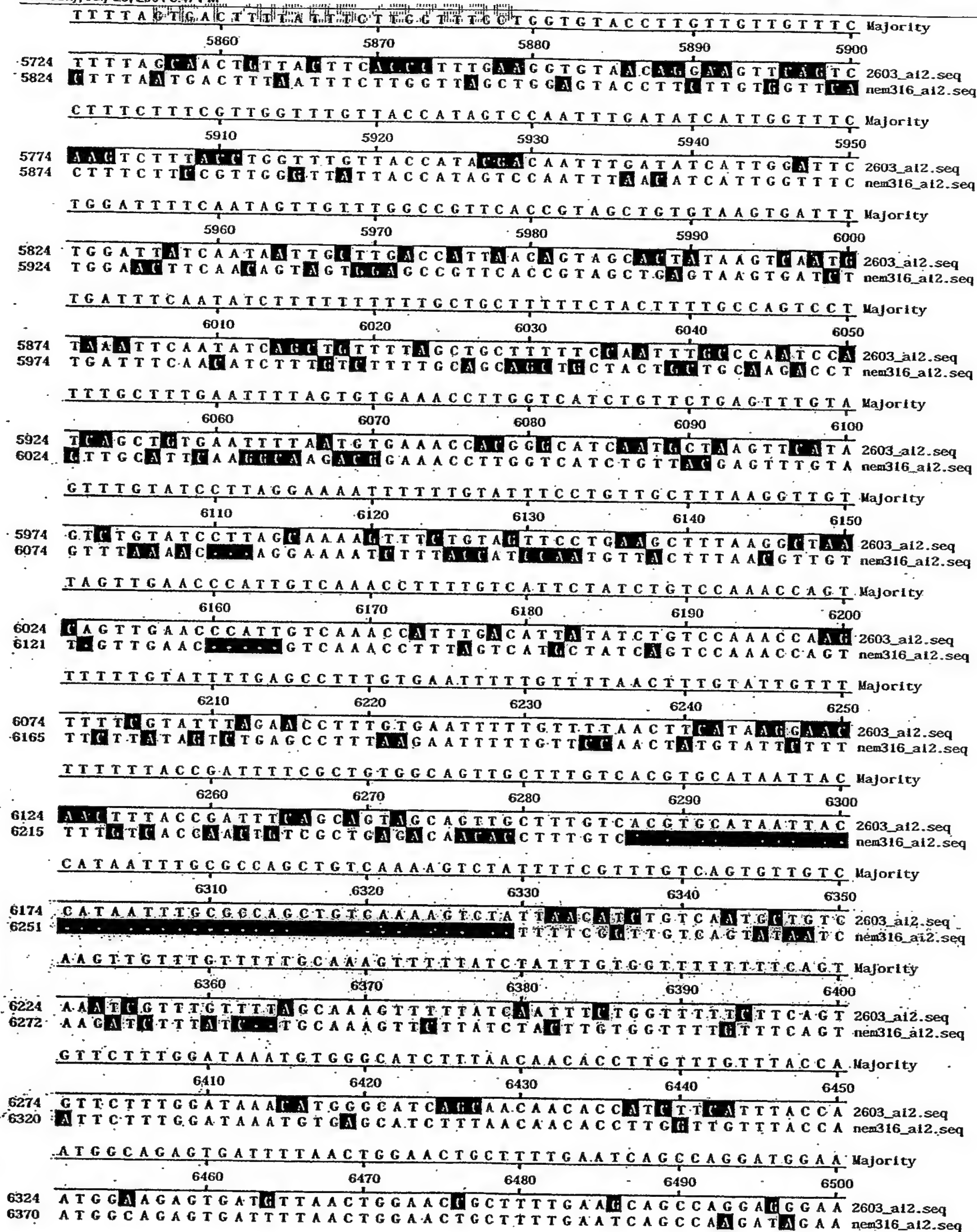


FIGURE 20I

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CCGTTTCTTCTGTTAGCTTTGATTTTCTTTCACTTCAACGATTTTCTACTT Majority
6510 6520 6530 6540 6550
6374 CCAATTATTGTTGTAAAGTAGATTTTCTTTTAACTTCAACAAATTTTAAACTT 2603_al2.seq
6420 CCGTTTGTATCTGTTAGTTTGAATTTTCTTTTCAATTCAACGATTTTGTAAAGT nem316_al2.seq

TCCTTTTTAATTTTTTGGTGTGTTGAAAGCAAGTCCAGTGTCTTTTTTTGGTG Majority
6560 6570 6580 6590 6600
6424 GCCTTTTAAATCTTTTGGTGTGTTGAAAACAAGTCCAGTATCTTCTCTCTGCTG 2603_al2.seq
6470 TCCTTTTTAATTTAGGAGTGTGTTAAAAGCAAACAAGTGTCTTTTTTTTAACT nem316_al2.seq

TTGATCCAGGCACGGCCTCATCTTTATTTTCTTTTGTGTTCCGGAGTATCG Majority
6610 6620 6630 6640 6650
6474 TCAATCCAGACACGGCCTCATCAATATTTACTGTTATTTTCAAGGAGTATCA 2603_al2.seq
6520 CTGA...AAGAACAGCAACCTTCAAGATCTTTAGCTTCCAAAGTATCG nem316_al2.seq

TCTTTCTTAATTAAGGCTGGTGTGTTAATTTGTTACCTTCTTTTTTCTTAAT Majority
6660 6670 6680 6690 6700
6524 TCTTTTAAATTAAGGCTGGTGTGTTAATTTGTTACCTTCTTTTGCCTTAAT 2603_al2.seq
6567 ACTTCTTA...CTTCTTAATTTTCAATTTTCAAGTAAAT nem316_al2.seq

GTATTGCATTTTACCAGTTTATTTTTTTTTTCAAAGCTAAAGCAAAGAACG Majority
6710 6720 6730 6740 6750
6574 ATATTGCACTTTTACCACTTTTATCTTTCTTTCAAAGCTAAAGCAAAGAACG 2603_al2.seq
6590 GATT...TTGTACCAGTTTCAATTTT...AAAACAAGAAAG nem316_al2.seq

CACCTTTGATTTCTTTAGCTTCGTTTGAGCCAAAGTAAGCTTTAAGGTCA Majority
6760 6770 6780 6790 6800
6624 CACCTTGGATTTCTTTAGATCTCT...GCCAAAGTAATCAGCAAGGTCA 2603_al2.seq
6628 CACCCTTAATTTCTTTAGCATCTGTTTGAGCCAAATAAGATTTAAGGTCA nem316_al2.seq

TTAATTTGTTTACCTTTGTAGTCTTTTTTCTTTCTTACCTTTTGTTCCTTG Majority
6810 6820 6830 6840 6850
6671 GAAATAGCTCTACCTTTGTAGTCTTTTCTGTTAAGACCTGTAGTTCCTTG 2603_al2.seq
6678 TTAATTTGTTTACC...ATAATCTGCTATCAATCTTACCTTTTGTACCTTG nem316_al2.seq

GGAGTTACTTTTGTGTTAAGTTTGTCTTGTGTTTGTGACAATCTTGTGCAAGG Majority
6860 6870 6880 6890 6900
6721 GAAGTTACTTTTGTGTTAAGATTTGATTCTGCTTTGCAAAATCTTGTGCAAG 2603_al2.seq
6728 AG...TAAGTTATCAAAATGCAAGCTTGTGCAATGACAATCTTGTGCAAGG nem316_al2.seq

TCACTGTATTAGTTGTTGCTTCTGTCGGCAAACGCTGGTGCAACTGAGAGT Majority
6910 6920 6930 6940 6950
6771 TCACTGTATTAGTTGTTGCTTCTATCCGCAAACGCTGGTGCAACTGAGAGT 2603_al2.seq
6775 TCACAGTATGAGTTGTTCTGTCGGCAAACGCTGGTGCAACTGAGAA nem316_al2.seq

AGTGACGTTAAGGTCAGTAGCAGTGTGCGAGAACATTGTAAGATATTTGTT Majority
6960 6970 6980 6990 7000
6821 ATGACGTTAAGGTCAGTAGCAATGCGAGAACATTGTAAGATATTTGTT 2603_al2.seq
6825 AGTGACGTTAAGGTCAGTAGCAGTGTGCGAGAACATTGTAAGATATTTGTT nem316_al2.seq

GATTTTTTTTCAATTTCTATCTCTTCTTATTTTAGTTAATCAACATGGTTA Majority
7010 7020 7030 7040 7050
6871 GATTCTTTTCAATTTCTATCTCTTCTTATTTTAGTTAATCAACATGATTA 2603_al2.seq
6875 GATTTTTTTTCAATTTCTATCTCTTCTTATTTTAGTTAATCAACATAGATA nem316_al2.seq

ATAATATGCGGATTTTAATATTACCGCAGGACCACTCCTTTCAAGTCAATG Majority
7060 7070 7080 7090 7100
6921 ATAATATGCGGATTTTAATATACCGCAGGACCACTCCTTTCAAGTCAATG 2603_al2.seq
6925 ATAATATGCGGATTTTAATATTACCGCAGGACCACTCCTTTT... nem316_al2.seq

GAATTTTTATTTAATTAATTAAGAATACTAAAGCGCATGATTTTTAATCTT Majority
7110 7120 7130 7140 7150
6970 GAATTTTTATTTAATTAATTAAGAATACTAAAGCGCATATTTTTAATCTT 2603_al2.seq
6967 GAATTTATTTAATTAATTAAGAATACTAAACACATGATTGTTAATCTT nem316_al2.seq
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FIGURE 20J



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	TTTTTCTGGATATATCACTAGATTTCCTTATATCTTTTCCAAATATAAAATT	Majority
	7160 7170 7180 7190 7200	
7020	TTTTGATGGATATATCACTAGATTTCCTTATAGCTTTTCCAAATATAAAATT	2603_al2.seq
7017	TTTTCTCTAGATATATCACTAGATTTCCTTATATCTTTTCCAAATATAAAATT	nem316_al2.seq
	CCACCTGCAATAGACATCATAGCTCCACCTATTAAAAATGAAAGATAGAAAT	Majority
	7210 7220 7230 7240 7250	
7070	CCACCTGCAATAGACATCATAGCTCCACCTATTAAAAATGAAAGATAGAAAT	2603_al2.seq
7067	CCACCTGCAATAGACATCATAGCTCCACCTATTAAAAATGAAAGATAGAAAT	nem316_al2.seq
	TCCTTTCCCACCTGTCATCGGAATAATTCCCTTTTGGTGGAAATATGCGTGT	Majority
	7260 7270 7280 7290 7300	
7120	TCCTTTCCCACCTGTCATCGGAATAATTCCCTTTTGGTGGAAATATGCGTGT	2603_al2.seq
7117	TCCTTTCCCACCTGTCATCGGAATAATTCCCTTTTGGTGGAAATATGCGTGT	nem316_al2.seq
	TGGTAATTAAATGCTTGTACCTTCCTCATGATATTCAGAAATCTGTTTA	Majority
	7310 7320 7330 7340 7350	
7170	TGGTAATTAAATGCTTGTACCTTCCTCATGATATTCAGAAATCTGTTTA	2603_al2.seq
7167	TGGTAATTAAATGCTTGTACCTTCCTCATGATATTCAGAAATCTGTTTA	nem316_al2.seq
	TTAACAGCTATTATATTTTTTATCGATCCTTTTAACCACTTCAAAAGTTAA	Majority
	7360 7370 7380 7390 7400	
7220	TTAACAGCTATTATATTTTTTATCGATCCTTTTAACCACTTCAAAAGTTAA	2603_al2.seq
7217	TTAACAGCTATTATATTTTTTATCGATCCTTTTAACCACTTCAAAAGTTAA	nem316_al2.seq
	AATTGGTTTATTAGTAATTTTTTGATAATCCTTCGGCGAAACTGCTTCTA	Majority
	7410 7420 7430 7440 7450	
7270	AATTGGTTTATTAGTAATTTTTTGATAATCCTTCGGCGAAACTGCTTCTA	2603_al2.seq
7267	AATTGGTTTATTAGTAATTTTTTGATAATCCTTCGGCGAAACTGCTTCTA	nem316_al2.seq
	TTAACTGATATTTGCCATCTTTCAAATCTTTGTAAGAAATTTTGCCGTTT	Majority
	7460 7470 7480 7490 7500	
7320	TTAACTGATATTTGCCATCTTTCAAATCTTTGTAAGAAATTTTGCCGTTT	2603_al2.seq
7317	TTAACTGATATTTGCCATCTTTCAAATCTTTGTAAGAAATTTTGCCGTTT	nem316_al2.seq
	TCTCCCGTCACTACTTTTGAATTATTATTTTTTATTGCTAAATAAAAGTTT	Majority
	7510 7520 7530 7540 7550	
7370	TCTCCCGTCACTACTTTTGAATTATTATTTTTTATTGCTAAATAAAAGTTT	2603_al2.seq
7367	TCTCCCGTCACTACTTTTGAATTATTATTTTTTATTGCTAAATAAAAGTTT	nem316_al2.seq
	ATAATCTTCATTAAATTCTTTGAAGTTCAAACGCTAGCTCCTTTGAGAAGCA	Majority
	7560 7570 7580 7590 7600	
7420	ATAATCTTCATTAAATTCTTTGAAGTTCAAACGCTAGCTCCTTTGAGAAGCA	2603_al2.seq
7417	ATAATCTTCATTAAATTCTTTGAAGTTCAAACGCTAGCTCCTTTGAGAAGCA	nem316_al2.seq
	ACTTATTATTATCTTTATCAACTTTTGTAAATTCAATTTCACCTAACTTC	Majority
	7610 7620 7630 7640 7650	
7470	ACTTATTATTATCTTTATCAACTTTTGTAAATTCAATTTCACCTAACTTC	2603_al2.seq
7467	ACTTATTATTATCTTTATCAACTTTTGTAAATTCAATTTCACCTAACTTC	nem316_al2.seq
	TTCTCGTTTTTAATCGTTATTGTAGGATATTCTCTCACATCAGGAATTTT	Majority
	7660 7670 7680 7690 7700	
7520	TTCTCGTTTTTAATCGTTATTGTAGGATATTCTCTCACATCAGGAATTTT	2603_al2.seq
7517	TTCTCGTTTTTAATCGTTATTGTAGGATATTCTCTCACATCAGGAATTTT	nem316_al2.seq
	AGGGATTGGAAAATCTCTAAGTGTATTAGGATCCTCTGATTTAGGATTCA	Majority
	7710 7720 7730 7740 7750	
7570	AGGGATTGGAAAATCTCTAAGTGTATTAGGATCCTCTGATTTAGGATTCA	2603_al2.seq
7567	AGGGATTGGAAAATCTCTAAGTGTATTAGGATCCTCTGATTTAGGATTCA	nem316_al2.seq
	ATGTTGTTCTACCATTAAGTGTATAGAAATTTGTTACTTATAAAACTGTCA	Majority
	7760 7770 7780 7790 7800	
7620	ATGTTGTTCTACCATTAAGTGTATAGAAATTTGTTACTTATAAAACTGTCA	2603_al2.seq
7617	ATGTTGTTCTACCATTAAGTGTATAGAAATTTGTTACTTATAAAACTGTCA	nem316_al2.seq

FIGURE 20K

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	TCTAGTTTTCACATCATATGTCAGTGTACTTTTTTGTCCCTTCTCCTAAGTT	Majority
	7810 7820 7830 7840 7850	
7670	TCTAGTTTTCACATCATATGTCAGTGTACTTTTTTGTCCCTTCTCCTAAGTT	2603_al2.seq
7667	TCTAGTTTTCACATCATATGTCAGTGTACTTTTTTGTCCCTTCTCCTAAGTT	nem316_al2.seq
	CAAACCTCTAACGTAGAGTTTATTTTTTGTATGTAATTCTAATTTAACCCTT	Majority
	7860 7870 7880 7890 7900	
7720	CAAACCTCTAACGTAGAGTTTATTTTTTGTATGTAATTCTAATTTAACCCTT	2603_al2.seq
7717	CAAACCTCTAACGTAGAGTTTATTTTTTGTATGTAATTCTAATTTAACCCTT	nem316_al2.seq
	TAAGTATTCCACCATCATTATTAGGCCCAACCAGTTGCAATGCTATCTTTT	Majority
	7910 7920 7930 7940 7950	
7770	TAAGTATTCCACCATCATTATTAGGCCCAACCAGTTGCAATGCTATCTTTT	2603_al2.seq
7767	TAAGTATTCCACCATCATTATTAGGCCCAACCAGTTGCAATGCTATCTTTT	nem316_al2.seq
	ATTATACTTCCATCATTTCCTGTAAAGTATAATCACTTGCTTGTAAATGT	Majority
	7960 7970 7980 7990 8000	
7820	ATTATACTTCCATCATTTCCTGTAAAGTATAATCACTTGCTTGTAAATGT	2603_al2.seq
7817	ATTATACTTCCATCATTTCCTGTAAAGTATAATCACTTGCTTGTAAATGT	nem316_al2.seq
	TTGTCCGTTGCCAAGCTGTAAATTGATTTTGTACCCCATAGGATCTTCTA	Majority
	8010 8020 8030 8040 8050	
7870	TTGTCCGTTGCCAAGCTGTAAATTGATTTTGTACCCCATAGGATCTTCTA	2603_al2.seq
7867	TTGTCCGTTGCCAAGCTGTAAATTGATTTTGTACCCCATAGGATCTTCTA	nem316_al2.seq
	FAGTTCCATTAAACAATTGAGTTTTCTTTTGTAAATCTTTTCAAATTGT	Majority
	8060 8070 8080 8090 8100	
7920	TAGTTCCATTAAACAATTGAGTTTTCTTTTGTAAATCTTTTCAAATTGT	2603_al2.seq
7917	TAGTTCCATTAAACAATTGAGTTTTCTTTTGTAAATCTTTTCAAATTGT	nem316_al2.seq
	TGCTGAATTTTAGATAAAATTTCAATTGTTAGATGTATCGGCTGAAGTTAC	Majority
	8110 8120 8130 8140 8150	
7970	TGCTGAATTTTAGATAAAATTTCAATTGTTAGATGTATCGGCTGAAGTTAC	2603_al2.seq
7967	TGCTGAATTTTAGATAAAATTTCAATTGTTAGATGTATCGGCTGAAGTTAC	nem316_al2.seq
	TATCGGGGTGTAGTACTCAGGTTTGGGAAGAGAATGACTTTCATTAGTTCTG	Majority
	8160 8170 8180 8190 8200	
8020	TATCGGGGTGTAGTACTCAGGTTTGGGAAGAGAATGACTTTCATTAGTTCTG	2603_al2.seq
8017	TATCGGGGTGTAGTACTCAGGTTTGGGAAGAGAATGACTTTCATTAGTTCTG	nem316_al2.seq
	TTATTTCTCCATCTGAAAGTTTAAAGCTTCTCTTTCAATTTTTTGAAAA	Majority
	8210 8220 8230 8240 8250	
8070	TTATTTCTCCATCTGAAAGTTTAAAGCTTCTCTTTCAATTTTTTGAAAA	2603_al2.seq
8067	TTATTTCTCCATCTGAAAGTTTAAAGCTTCTCTTTCAATTTTTTGAAAA	nem316_al2.seq
	GTACCATCTTGATTTTTTCTTATACTCCTCATTATAAACTTGTCTAAAAGC	Majority
	8260 8270 8280 8290 8300	
8120	GTACCATCTTGATTTTTTCTTATACTCCTCATTATAAACTTGTCTAAAAGC	2603_al2.seq
8117	GTACCATCTTGATTTTTTCTTATACTCCTCATTATAAACTTGTCTAAAAGC	nem316_al2.seq
	AGATATATCTATACCAAAATTAAGATGTCATAATTTTCTGTTTTTAAAC	Majority
	8310 8320 8330 8340 8350	
8170	AGATATATCTATACCAAAATTAAGATGTCATAATTTTCTGTTTTTAAAC	2603_al2.seq
8167	AGATATATCTATACCAAAATTAAGATGTCATAATTTTCTGTTTTTAAAC	nem316_al2.seq
	TATTTATATAAAGTTTGGTTGGTGTTCATGTTCTTTTACTGGTCCATTT	Majority
	8360 8370 8380 8390 8400	
8220	TATTTATATAAAGTTTGGTTGGTGTTCATGTTCTTTTACTGGTCCATTT	2603_al2.seq
8217	TATTTATATAAAGTTTGGTTGGTGTTCATGTTCTTTTACTGGTCCATTT	nem316_al2.seq
	CGATAAATTGTACCTTTAGGGTAATTAAGATTAAATCTAAATAATGAAG	Majority
	8410 8420 8430 8440 8450	
8270	CGATAAATTGTACCTTTAGGGTAATTAAGATTAAATCTAAATAATGAAG	2603_al2.seq
8267	CGATAAATTGTACCTTTAGGGTAATTAAGATTAAATCTAAATAATGAAG	nem316_al2.seq

FIGURE 20L



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	TTTTTGTAAAGTTTCCAGAGATTATCTGTGTTTGTATAACTATCTAAGGGAA	Majority
	8460 8470 8480 8490 8500	
8320	TTTTTGTAAAGTTTCCAGAGATTATCTGTGTTTGTATAACTATCTAAGGGAA	2603_al2.seq
8317	TTTTTGTAAAGTTTCCAGAGATTATCTGTGTTTGTATAACTATCTAAGGGAA	nem316_al2.seq
	ACAAAAAGTAACCTCTCCCAATTTCTTTTATATCCTCGGGCTTATCAGTA	Majority
	8510 8520 8530 8540 8550	
8370	ACAAAAAGTAACCTCTCCCAATTTCTTTTATATCCTCGGGCTTATCAGTA	2603_al2.seq
8367	ACAAAAAGTAACCTCTCCCAATTTCTTTTATATCCTCGGGCTTATCAGTA	nem316_al2.seq
	AGTAGAAAATTACTTTTATTTAGATATCCATTTTTTTTTCATTTGTTCAA	Majority
	8560 8570 8580 8590 8600	
8420	AGTAGAAAATTACTTTTATTTAGATATCCATTTTTTTTTCATTTGTTCAA	2603_al2.seq
8417	AGTAGAAAATTACTTTTATTTAGATATCCATTTTTTTTTCATTTGTTCAA	nem316_al2.seq
	TTGGCTTTTCATATGATGCACCCAGTTTAAATTATTAATAGCATATGATC	Majority
	8610 8620 8630 8640 8650	
8470	TTGGCTTTTCATATGATGCACCCAGTTTAAATTATTAATAGCATATGATC	2603_al2.seq
8467	TTGGCTTTTCATATGATGCACCCAGTTTAAATTATTAATAGCATATGATC	nem316_al2.seq
	TTGTTGGAACACCATCAGTTATATGAACAATAATTTTTTGAATTTTCGA	Majority
	8660 8670 8680 8690 8700	
8520	TTGTTGGAACACCATCAGTTATATGAACAATAATTTTTTGAATTTTCGA	2603_al2.seq
8517	TTGTTGGAACACCATCAGTTATATGAACAATAATTTTTTGAATTTTCGA	nem316_al2.seq
	TTTACTTGAATCAAAATATCATCTGCTCCATGAAGGCTTTTCATAGTAAA	Majority
	8710 8720 8730 8740 8750	
8570	TTTACTTGAATCAAAATATCATCTGCTCCATGAAGGCTTTTCATAGTAAA	2603_al2.seq
8567	TTTACTTGAATCAAAATATCATCTGCTCCATGAAGGCTTTTCATAGTAAA	nem316_al2.seq
	TGTTTCTCCTACTTTACTAAGATAGTACTGCTTTTGTGCTCTGGAGTTA	Majority
	8760 8770 8780 8790 8800	
8620	TGTTTCTCCTACTTTACTAAGATAGTACTGCTTTTGTGCTCTGGAGTTA	2603_al2.seq
8617	TGTTTCTCCTACTTTACTAAGATAGTACTGCTTTTGTGCTCTGGAGTTA	nem316_al2.seq
	GTCCGTTTGTAGTTGATCCCAATTTAGCTTTTAGGAGCTTCTGTCCGAAATC	Majority
	8810 8820 8830 8840 8850	
8670	GTCCGTTTGTAGTTGATCCCAATTTAGCTTTTAGGAGCTTCTGTCCGAAATC	2603_al2.seq
8667	GTCCGTTTGTAGTTGATCCCAATTTAGCTTTTAGGAGCTTCTGTCCGAAATC	nem316_al2.seq
	CTTTTTTATAATCTCTTCAGCATTATTTTGTAAATTGTTTATGACTATAATT	Majority
	8860 8870 8880 8890 8900	
8720	CTTTTTTATAATCTCTTCAGCATTATTTTGTAAATTGTTTATGACTATAATT	2603_al2.seq
8717	CTTTTTTATAATCTCTTCAGCATTATTTTGTAAATTGTTTATGACTATAATT	nem316_al2.seq
	CTCTGTCTGAATTGTGAACCTTAGTTTGAAGGCCATAAATATTTATCATCTT	Majority
	8910 8920 8930 8940 8950	
8770	CTCTGTCTGAATTGTGAACCTTAGTTTGAAGGCCATAAATATTTATCATCTT	2603_al2.seq
8767	CTCTGTCTGAATTGTGAACCTTAGTTTGAAGGCCATAAATATTTATCATCTT	nem316_al2.seq
	CTTTAAATCCCTTTTACGACATCTACACTCCTACCATCAAAAATATCTGAA	Majority
	8960 8970 8980 8990 9000	
8820	CTTTAAATCCCTTTTACGACATCTACACTCCTACCATCAAAAATATCTGAA	2603_al2.seq
8817	CTTTAAATCCCTTTTACGACATCTACACTCCTACCATCAAAAATATCTGAA	nem316_al2.seq
	CCATAGGTAACATAATGCAACCCTATTATCACTGTTTGTCTCCTAAAAATATC	Majority
	9010 9020 9030 9040 9050	
8870	CCATAGGTAACATAATGCAACCCTATTATCACTGTTTGTCTCCTAAAAATATC	2603_al2.seq
8867	CCATAGGTAACATAATGCAACCCTATTATCACTGTTTGTCTCCTAAAAATATC	nem316_al2.seq
	TTTTACTGCGGTCCCAAGAGCTTCGGCAGCTTTCTTGGCTTTATTATGCC	Majority
	9060 9070 9080 9090 9100	
8920	TTTTACTGCGGTCCCAAGAGCTTCGGCAGCTTTCTTGGCTTTATTATGCC	2603_al2.seq
8917	TTTTACTGCGGTCCCAAGAGCTTCGGCAGCTTTCTTGGCTTTATTATGCC	nem316_al2.seq

FIGURE 20M

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	TTTGA	AAATTT	TGGGCC	ATCGTT	ATTTCATT	GAGTTAG	AATTAT	CGAGTAC	G	Majority
	9110		9120		9130		9140		9150	
8970	TTTGA	AAATTT	TGGGCC	ATCGTT	ATTTCATT	GAGTTAG	AATTAT	CGAGTAC	G	2603_al2.seq
8967	TTTGA	AAATTT	TGGGCC	ATCGTT	ATTTCATT	GAGTTAG	AATTAT	CGAGTAC	G	nem316_al2.seq
	AAGACAACATCTAACGGCCTTTTGTGTTGTCCACTGCTTTTACTATGGTTTTT									
	9160		9170		9180		9190		9200	
9020	AAGACA	ACATCT	AACGGC	CTTTTGT	GTTGTCC	ACTGCT	TTTTTACT	ATGGTTTTT		2603_al2.seq
9017	AAGACA	ACATCT	AACGGC	CTTTTGT	GTTGTCC	ACTGCT	TTTTTACT	ATGGTTTTT		nem316_al2.seq
	TCCACTGACAGTTAACTCAATTTTATATTTATTATGAGCTAAATCACCTA									
	9210		9220		9230		9240		9250	
9070	TCCACT	GACAGT	TAACTCA	ATTTTAT	ATTTATT	ATGAGCT	AAATCAC	CTA		2603_al2.seq
9067	TCCACT	GACAGT	TAACTCA	ATTTTAT	ATTTATT	ATGAGCT	AAATCAC	CTA		nem316_al2.seq
	CTTCTGAAATACGTTTATAGATAATGTTCCCTCTGGAATTTCTCTTATATGC									
	9260		9270		9280		9290		9300	
9120	CTTCTG	AAATAC	GTTTATAG	ATAATG	TTCCTCT	GGAATTT	CTCTT	ATATGC		2603_al2.seq
9117	CTTCTG	AAATAC	GTTTATAG	ATAATG	TTCCTCT	GGAATTT	CTCTT	ATATGC		nem316_al2.seq
	TCACCTTCACTTGAATATGGGTTAACTGCTTTTGCCTCTGACTTTCCATT									
	9310		9320		9330		9340		9350	
9170	TCACCT	TCACTT	GAAATAT	GGGTTAA	CTGCTTT	TGCCTCT	GACTTT	CCATT		2603_al2.seq
9167	TCACCT	TCACTT	GAAATAT	GGGTTAA	CTGCTTT	TGCCTCT	GACTTT	CCATT		nem316_al2.seq
	TGGAACTGAACCTTTAACAATGCTCAAGTTTATAAGATTCTTTGTATCTT									
	9360		9370		9380		9390		9400	
9220	TGGAACT	GAACCT	TTAACA	ATGCTCA	AGTTTAT	AAGATT	CTTTGT	ATCTT		2603_al2.seq
9217	TGGAACT	GAACCT	TTAACA	ATGCTCA	AGTTTAT	AAGATT	CTTTGT	ATCTT		nem316_al2.seq
	CATAAATTCTCTGTGGGGGGATACTGCTTATCTAGTTCTTCTGATTTTGT									
	9410		9420		9430		9440		9450	
9270	CATAA	ATTCTC	TGTGGG	GGGATA	CTGCTT	ATCTAG	TTCTTCT	GATTTTGT		2603_al2.seq
9267	CATAA	ATTCTC	TGTGGG	GGGATA	CTGCTT	ATCTAG	TTCTTCT	GATTTTGT		nem316_al2.seq
	CCAATTGTGGAATTTTATCAACCACTATTTTGTATCGTAGTTTTTCCATT									
	9460		9470		9480		9490		9500	
9320	CCAATT	GTGGA	ATTTTAT	CAACCA	CTATTT	TGTATCG	TAGTTT	TCCATT		2603_al2.seq
9317	CCAATT	GTGGA	ATTTTAT	CAACCA	CTATTT	TGTATCG	TAGTTT	TCCATT		nem316_al2.seq
	ACTCTCAACCTTAACTTGCCAAAGTCTGGTTAGTCTTTTTTATAACCTTCGG									
	9510		9520		9530		9540		9550	
9370	ACTCTC	AACCTT	AACTTG	CCAAAG	TCTGGT	TAGTCT	TTTTTAT	AACCTTC	GG	2603_al2.seq
9367	ACTCTC	AACCTT	AACTTG	CCAAAG	TCTGGT	TAGTCT	TTTTTAT	AACCTTC	GG	nem316_al2.seq
	GCGCTGTTTCTTCTGATAAAGTATAATCTCCAGGTATGAGATTATCAAAA									
	9560		9570		9580		9590		9600	
9420	GCGCTG	TTTCTT	CTGATA	AAGTATA	ATCTCC	AGGTAT	GAGATT	ATCAAAA		2603_al2.seq
9417	GCGCTG	TTTCTT	CTGATA	AAGTATA	ATCTCC	AGGTAT	GAGATT	ATCAAAA		nem316_al2.seq
	GTAGCTTCACTGTTAGCTCAGCAGTTACTTTTTCTATTTTACTTTCTGG									
	9610		9620		9630		9640		9650	
9470	GTAGCT	TCACTG	TTAGCT	CAGCAG	TTACTT	TTTTCT	ATTTTAC	TTTCTGG		2603_al2.seq
9467	GTAGCT	TCACTG	TTAGCT	CAGCAG	TTACTT	TTTTCT	ATTTTAC	TTTCTGG		nem316_al2.seq
	ATGAGCAGTAGTTTTTAAACAAAGGTAGCTTTTGAAAGTGGTTTCTTCT									
	9660		9670		9680		9690		9700	
9520	ATGAGC	AGTAGT	TTTTTAA	ACAAAG	GTAGCT	TTTGAA	AGTGGT	TTTCTTCT		2603_al2.seq
9517	ATGAGC	AGTAGT	TTTTTAA	ACAAAG	GTAGCT	TTTGAA	AGTGGT	TTTCTTCT		nem316_al2.seq
	GGTCATCTGTCTTTTTTAAACAACCTAATTTCTTTAGCACCATTTTCCGGT									
	9710		9720		9730		9740		9750	
9570	GGTCAT	CTGTCT	TTTTTAA	ACAAC	CTAAT	TTCTTT	AGCAC	CATTTT	CCGGT	2603_al2.seq
9567	GGTCAT	CTGTCT	TTTTTAA	ACAAC	CTAAT	TTCTTT	AGCAC	CATTTT	CCGGT	nem316_al2.seq

FIGURE 20N



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ACGGTACCTTTCCCTAAACATTTGGTATTAAAGCGGTATTTGCGACAAACA Majority  
9760 9770 9780 9790 9800  
9620 ACGGTACTTTCCCTAAACATTTGGTATTAAAGCGGTATTTGCGACAAACA 2603\_al2.seq  
9617 ACGGTACTTTCCCTAAACATTTGGTATTAAAGCGGTATTTGCGACAAACA nem316\_al2.seq  
AAAAAGACTTAAACGTCAATATTTTAGAAAATTTTGGTATTTTCTCATTT Majority  
9810 9820 9830 9840 9850  
9670 AAAAAGACTTAAACGTCAATATTTTAGAAAATTTTGGTATTTTCTCATTT 2603\_al2.seq  
9667 AAAAAGACTTAAACGTCAATATTTTAGAAAATTTTGGTATTTTCTCATTT nem316\_al2.seq  
TACAACCTCCTATTGTGCGCGAAATGTCGTTTCTAAATCTAAGATCAGATAC Majority  
9860 9870 9880 9890 9900  
9720 TACAACCTCCTATTGTGCGCGAAATGTCGTTTCTAAATCTAAGATCAGATAC 2603\_al2.seq  
9717 TACAACCTCCTATTGTGCGCGAAATGTCGTTTCTAAATCTAAGATCAGATAC nem316\_al2.seq  
AGAATATCCTAGAAATATACAAACTATCACTTATTATGATATCAATAATTT Majority  
9910 9920 9930 9940 9950  
9770 AGAATATCCTAGAAATATACAAACTATCACTTATTATGATATCAATAATTT 2603\_al2.seq  
9767 AGAATATCCTAGAAATATACAAACTATCACTTATTATGATATCAATAATTT nem316\_al2.seq  
CTTATTATAAGGTATGGAATTTTAATGTTTTTTCCCAATTTTGAATGAT Majority  
9960 9970 9980 9990 10000  
9820 CTTATTATAAGGTATGGAATTTTAATGTTTTTTCCCAATTTTGAATGAT 2603\_al2.seq  
9817 CTTATTATAAGGTATGGAATTTTAATGTTTTTTCCCAATTTTGAATGAT nem316\_al2.seq  
TTTTCTTTTTTATTGATAATCTTATTTTTTTATTATCTTAGAAATATTTCA Majority  
10010 10020 10030 10040 10050  
9870 TTTTCTTTTTTATTGATAATCTTATTTTTTTATTATCTTAGAAATATTTCA 2603\_al2.seq  
9867 TTTTCTTTTTTATTGATAATCTTATTTTTTTATTATCTTAGAAATATTTCA nem316\_al2.seq  
ATTAGCTTAAGTAGTTGATTTTTCTTTTTTTATGTTTTTAAATATTGCTT Majority  
10060 10070 10080 10090 10100  
9920 ATTAGCTTAAGTAGTTGATTTTTCTTTTTTTATGTTTTTAAATATTGCTT 2603\_al2.seq  
9917 ATTAGCTTAAGTAGTTGATTTTTCTTTTTTTATGTTTTTAAATATTGCTT nem316\_al2.seq  
AAAAATAATGTTTGAGAGAGAGTTTACTGAATTGATTGAAAATTATTTAG Majority  
10110 10120 10130 10140 10150  
9970 AAAAATAATGTTTGAGAGAGAGTTTACTGAATTGATTGAAAATTATTTAG 2603\_al2.seq  
9967 AAAAATAATGTTTGAGAGAGAGTTTACTGAATTGATTGAAAATTATTTAG nem316\_al2.seq  
AAAAAGACATCCTTAATCAAAATAAAACTTCTAACTTTATGCTATGATTAC Majority  
10160 10170 10180 10190 10200  
10018 AAAAAGACATCCTTAATCAAAATAAAACTTCTAACTTTATGCTATGATTAC 2603\_al2.seq  
10017 AAAAAGACATCCTTAATCAAAATAAAACTTCTAACTTTATGCTATGATTAC nem316\_al2.seq  
TACCCTTCCATTACTCTAGACAAATCATGTCATCAACTTGGTTTATCTGA Majority  
10210 10220 10230 10240 10250  
10068 TACCCTTCCATTACTCTAGACAAATCATGTCATCAACTTGGTTTATCTGA 2603\_al2.seq  
10067 TACCCTTCCATTACTCTAGACAAATCATGTCATCAACTTGGTTTATCTGA nem316\_al2.seq  
ACTACTTATTAGGAAATATTGTCATGATTTAACAACCTTTATTTAACAGTC Majority  
10260 10270 10280 10290 10300  
10118 ACTACTTATTAGGAAATATTGTCATGATTTAACAACCTTTATTTAACAGTC 2603\_al2.seq  
10117 ACTACTTATTAGGAAATATTGTCATGATTTAACAACCTTTATTTAACAGTC nem316\_al2.seq  
AACTCTCTCTGAATATCGAAAAGAGTACAATAGTCTACCAATCTAATGGT Majority  
10310 10320 10330 10340 10350  
10168 AACTCTCTCTGAATATCGAAAAGAGTACAATAGTCTACCAATCTAATGGT 2603\_al2.seq  
10167 AACTCTCTCTGAATATCGAAAAGAGTACAATAGTCTACCAATCTAATGGT nem316\_al2.seq  
GTAACCTAGAGAACAAAGCTTTCAAATATATTTATCATCAATCACACGTTTT Majority  
10360 10370 10380 10390 10400  
10218 GTAACCTAGAGAACAAAGCTTTCAAATATATTTATCATCAATCACACGTTTT 2603\_al2.seq  
10217 GTAACCTAGAGAACAAAGCTTTCAAATATATTTATCATCAATCACACGTTTT nem316\_al2.seq

FIGURE 200

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	ACAACTTTTAAATTTTGTGATTCAGGCAGGTTACCTTTAA	Majority
	10410 10420 10430 10440 10450	
10268	ACAACTTTTAAATTTTGTGATTCAGGCAGGTTACCTTTAA	2603_al2.seq
10267	ACAACTTTTAAATTTTGTGATTCAGGCAGGTTACCTTTAA	nen316_al2.seq
	CTTACTTTAGTGAAAAATTTGGACTATCTTGTGCAACTGCTTATCGCATA	Majority
	10460 10470 10480 10490 10500	
10318	CTTACTTTAGTGAAAAATTTGGACTATCTTGTGCAACTGCTTATCGCATA	2603_al2.seq
10317	CTTACTTTAGTGAAAAATTTGGACTATCTTGTGCAACTGCTTATCGCATA	nen316_al2.seq
	CGAAAAACATATTAGTCCGTTACTAGAAAAACTTGGATTTCAGATTTTCAA	Majority
	10510 10520 10530 10540 10550	
10368	CGAAAAACATATTAGTCCATTACTAGAAAAACTTGGATTTCAGATTTTCAA	2603_al2.seq
10367	CGAAAAACATATTAGTCCGTTACTAGAAAAACTTGGATTTCAGATTTTCAA	nen316_al2.seq
	AAATACTATTACCGGTGACGAGTATCGAATTCGCTATTTAATCGCATTTT	Majority
	10560 10570 10580 10590 10600	
10418	AAATACTATTACCGGTGACGAGTATCGAATTCGCTATTTAATCGCATTTT	2603_al2.seq
10417	AAATACTATTACCGGTGACGAGTATCGAATTCGCTATTTAATCGCATTTT	nen316_al2.seq
	TAAATGCTCGATTGTTATAGAAAGTTTATCCCTTGTCTAAGATGGATAAA	Majority
	10610 10620 10630 10640 10650	
10468	TAAATGCTCGATTGTTATAGAAAGTTTATCCCTTGTCTAAGATGGATAAA	2603_al2.seq
10467	TAAATGCTCGATTGTTATAGAAAGTTTATCCCTTGTCTAAGATGGATAAA	nen316_al2.seq
	TTGCTTATCAAACGATTGTTATTAGAATACTCAACTACTTTTACTGCTTC	Majority
	10660 10670 10680 10690 10700	
10518	TTGCTTATCAAACGATTGTTATTAGAATACTCAACTACTTTTACTGCTTC	2603_al2.seq
10517	TTGCTTATCAAACGATTGTTATTAGAATACTCAACTACTTTTACTGCTTC	nen316_al2.seq
	TCATTACTTCCCAAATACATTTATTTTCTTTGATACATTGTTGTCTCTAT	Majority
	10710 10720 10730 10740 10750	
10568	TCATTACTTCCCAAATACATTTATTTTCTTTGATACATTGTTGTCTCTAT	2603_al2.seq
10567	TCATTACTTCCCAAATACATTTATTTTCTTTGATACATTGTTGTCTCTAT	nen316_al2.seq
	CATGGAAACGTATTAATTATAATGTAGTTGTCCCTTACTCATCCCTTTTC	Majority
	10760 10770 10780 10790 10800	
10618	CATGGAAACGTATTAATTATAATGTAGTTGTCCCTTACTCATCCCTTTTC	2603_al2.seq
10617	CATGGAAACGTATTAATTATAATGTAGTTGTCCCTTACTCATCCCTTTTC	nen316_al2.seq
	ATTGAACTACAAAAATATATTTATCTATGATACATTACAATATTGTGTCAA	Majority
	10810 10820 10830 10840 10850	
10668	ATTGAACTACAAAAATATATTTATCTATGATACATTACAATATTGTGTCAA	2603_al2.seq
10667	ATTGAACTACAAAAATATATTTATCTATGATACATTACAATATTGTGTCAA	nen316_al2.seq
	AAATGTTATTATAGATTCCCTTTAAATTTAATTTAAAAAAGACGATATAG	Majority
	10860 10870 10880 10890 10900	
10718	AAATGTTATTATAGATTCCCTTTAAATTTAATTTAAAAAAGACGATATAG	2603_al2.seq
10717	AAATGTTATTATAGATTCCCTTTAAATTTAATTTAAAAAAGACGATATAG	nen316_al2.seq
	ACTATATTTTCTTGTCTTACGTTACTTCCCATAACTCTTTTCTAATCCA	Majority
	10910 10920 10930 10940 10950	
10768	ACTATATTTTCTTGTCTTACGTTACTTCCCATAACTCTTTTCTAATCCA	2603_al2.seq
10767	ACTATATTTTCTTGTCTTACGTTACTTCCCATAACTCTTTTCTAATCCA	nen316_al2.seq
	AATTGGACTGAGAAGCGTATCGATAATGTAATAGCTATTTTCGAAAATTA	Majority
	10960 10970 10980 10990 11000	
10818	AATTGGACTGAGAAGCGTATCGATAATGTAATAGCTATTTTCGAAAATTA	2603_al2.seq
10817	AATTGGACTGAGAAGCGTATCGATAATGTAATAGCTATTTTCGAAAATTA	nen316_al2.seq
	TCCCAAATTCGAAAAATTTATTACAGCCACTCAAAGATGCTCTTCCCTTAT	Majority
	11010 11020 11030 11040 11050	
10868	TCCCAAATTCGAAAAATTTATTACAGCCACTCAAAGATGCTCTTCCCTTAT	2603_al2.seq
10867	TCCCAAATTCGAAAAATTTATTACAGCCACTCAAAGATGCTCTTCCCTTAT	nen316_al2.seq

FIGURE 20P



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CTGGCTCCTATCATGATGAGCTTCTGTTAAAGTTGCTATCATTITTTTCCGA Majority
11060 11070 11080 11090 11100
10918 CTGGCTCCTATCATGATGAGTTGGTAAAAGTTGCTATC TTTTITTTTCCGA 2603_al2.seq
10917 CTG A CTCCTATCATGA GAGTTGGTAAAAGTTGCTATCATTITTTT CCGA nem316_al2.seq

ACATTTATTTTAGGATTAAATCAATTAATCCCTGAAACAATTTTCATTTC Majority
11110 11120 11130 11140 11150
10967 ACATTTATTTTAGGATTAAATCAATTAATCCCTGAAACAATTTTCATTTC 2603_al2.seq
10967 ACATTTATTTTAGGATTAAATCAATTAATCCCTGAAACAATTTTCATTTC nem316_al2.seq

TTCATGGAACATATCATAGACATGATAAATTAACACTATTCTCCGACCGA Majority
11160 11170 11180 11190 11200
11017 TTCATGGAACATATCATAGACATGATAAATTAACACTATTCTCCGACCGA 2603_al2.seq
11017 TTCATGGAACATATCATAGACATGATAAATTAACACTATTCTCCGACCGA nem316_al2.seq

TAATTACAAATTGCTTAAGTGAAATTGGAGAATACACGTTTAAGGAACAA Majority
11210 11220 11230 11240 11250
11067 TAATTACAAATTGCTTAAGT A AATTGGAGAATACACGTTTA A GAACAA 2603_al2.seq
11067 TAATTACAAATTGCTTAAGTGAAATTGGAGAATACACGTTTAAGGAACAA nem316_al2.seq

CATTTTCTTCTCCTTTTGCTCATCTAGAAAGAATTATCAAAAATCATAT Majority
11260 11270 11280 11290 11300
11117 CATTTTCTTCTCCTTTTGTA CTCTCATCTAGAAAGAAT C ATCAAAAATCATAT 2603_al2.seq
11117 CATTTTCTTCTCCTTTTGCTCA CTAGAAAGAATTATCAAAAATCA A AT nem316_al2.seq

TCCTCCGATACAGATAGCCGTACTAACTACAGACTTTTATTAATAACCAA Majority
11310 11320 11330 11340 11350
11167 TCCTCCGATACAGATAGCCGTACTAACTACAGACTTTTATTAATAACCAA 2603_al2.seq
11167 TCCTCCGATACAA A ATAGCCGTACTAACTACAGACTTTTATTAATAAC A AAA nem316_al2.seq

TTTTAACAGAAATGTTTATTACAGAGGTTTCTTCTAAACAGATTTCATTTC Majority
11360 11370 11380 11390 11400
11217 TTTTAACAGAAATG T TATTACAGAG A TTTTCTTCTAA A AGATTTCATTTC 2603_al2.seq
11217 TTTTAACAGAAATGTTTAT C ACAGAGGTTTCTTCTTCTAAACAGATTTCATTTC nem316_al2.seq

CACCTTACTATTTATTAACCTGATGATCTTTCCAATATTACTAATCTTAA Majority
11410 11420 11430 11440 11450
11267 CACCTTACTATTTATTAACCTGATGATCTTTCCAATATTACTAATCTTAA 2603_al2.seq
11267 CACCTTACTATTTATTAACCTGATGATCTTTCCAATATTACTAATCTTAA nem316_al2.seq

TCCAGATATTATTATTACCAATCCAAAGCTTTCTTCTTTATCAAACATG Majority
11460 11470 11480 11490 11500
11317 TCCAGAA ATTATTATTACCAAT A AAGCTTTCT C CTTTATCAAACATG 2603_al2.seq
11317 TCCAGATATTATTATTACCA A CCAAAGCTTTCTTCTTTAT A AAACATG nem316_al2.seq

AGATTTCTTCAGAGAGTTTAATTACATATATTGATTGATTTCATACTTCA Majority
11510 11520 11530 11540 11550
11367 AGATTTCTTCAGAGAGTTT AATTACATATATTGAT T TGATT A TACTTCA 2603_al2.seq
11367 AGATTTCTTCAGAGAGTTT AATTACATATATTGATTGATTTCATACT C A nem316_al2.seq

GACCAGATCAATCAAAATCCAAGAAATTATTTTCATCAATACAGGAAGAAA Majority
11560 11570 11580 11590 11600
11417 GACCA A ATCAATCAAAATCCA A AATTATTTTCATCAATACAGGAAGAAA 2603_al2.seq
11417 GACCAGATCAATCAAAATCCAAGAAATTATTTTCATCAATAC A GAAGAAA nem316_al2.seq

ATATTGTAAACTTTTTTGCAAAAACATAATGAAATAACTACTCGTAGCTCCT Majority
11610 11620 11630 11640 11650
11467 ATATTG A AACTTTTTTGCAAAAACATAATGAAATAACT A TCGTAGCTCCT 2603_al2.seq
11467 ATATTGTAAACTTTTTTGCAAAA A AATGAAATAACTAC C TCGTAGCTCCT nem316_al2.seq

ATAACTCTTAAAAAATTAACATTA AAAAGCTAGAGCATTGTGTAATGCTCT Majority
11660 11670 11680 11690 11700
11517 ATA ACTCTTAAAAAATTAACATTA AAAAGCTAGAGCATTGTGTAATGCTCT 2603_al2.seq
11517 ATA ACTCTTAAAAAATTAACATTA AAAAGCTAGAGCATTGTGTAATGCTCT nem316_al2.seq

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FIGURE 20Q

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AGCTTTTAAATGTTAAATTTTGAATAATAATCCAACCTTTTCAACTG Majority  
11710 11720 11730 11740 11750  
11567 AGCTTTTAAATGTTAAATTTTGAATAATAATCCAACCTTTTCAACTG 2603\_al2.seq  
11567 AGCTTTTAAATGTTAAATTTTGAATAATAATCCAACCTTTTCAACTG nem316\_al2.seq  
TTTTTCCCATGTGAAATGTTCTTTAATTCTTTTAGCAATATTCTGTTGT Majority  
11760 11770 11780 11790 11800  
11617 TTTTTCCCATGTGAAATGTTCTTTAATTCTTTTAGCAATATTCTGTTGT 2603\_al2.seq  
11617 TTTTTCCCATGTGAAATGTTCTTTAATTCTTTTAGCAATATTCTGTTGT nem316\_al2.seq  
AGTTTCTCTCTTAATGCCTTATCTTTTACTAATAAATCAAGAGATTTCATG Majority  
11810 11820 11830 11840 11850  
11667 AGTTTCTCTCTTAATGCCTTATCTTTTACTAATAAATCAAGAGATTTCATG 2603\_al2.seq  
11667 AGTTTCTCTCTTAATGCCTTATCTTTTACTAATAAATCAAGAGATTTCATG nem316\_al2.seq  
GAGTGACTGAGTATTTTCTTCCATGATGATTCTTAACCTCAGGGCTATCAA Majority  
11860 11870 11880 11890 11900  
11717 GAGTGACTGAGTATTTTCTTCCATGATGATTCTTAACCTCAGGGCTATCAA 2603\_al2.seq  
11717 GAGTGACTGAGTATTTTCTTCCATGATGATTCTTAACCTCAGGGCTATCAA nem316\_al2.seq  
TAACTTCAACTGTTCCACCGCGATCTGTTGCAATAATAGCACTTGAAAGT Majority  
11910 11920 11930 11940 11950  
11767 TAACTTCAACTGTTCCACCGCGATCTGTTGCAATAATAGCACTTGAAAGT 2603\_al2.seq  
11767 TAACTTCAACTGTTCCACCGCGATCTGTTGCAATAATAGCACTTGAAAGT nem316\_al2.seq  
AGACCAGCTTCTAATAATAGAGGTTGGTAATCCCTCTGGATACATTGAAGG Majority  
11960 11970 11980 11990 12000  
11817 AGACCAGCTTCTAATAATAGAGGTTGGTAATCCCTCTGGATACATTGAAGG 2603\_al2.seq  
11817 AGACCAGCTTCTAATAATAGAGGTTGGTAATCCCTCTGGATACATTGAAGG nem316\_al2.seq  
GTAAACAAAGATATCAGTCTGTGCCATTAAAGACATAGTCTGTTCAAAGT Majority  
12010 12020 12030 12040 12050  
11867 GTAAACAAAGATATCAGTCTGTGCCATTAAAGACATAGTCTGTTCAAAGT 2603\_al2.seq  
11867 GTAAACAAAGATATCAGTCTGTGCCATTAAAGACATAGTCTGTTCAAAGT nem316\_al2.seq  
TTAATTTCCCAAAGTTAATCTGTTTGGACTGATATTTCTCTTTCAA Majority  
12060 12070 12080 12090 12100  
11917 TTAATTTCCCAAAGTTAATCTGTTTGGACTGATATTTCTCTTTCAA 2603\_al2.seq  
11917 TTAATTTCCCAAAGTTAATCTGTTTGGACTGATATTTCTCTTTCAA nem316\_al2.seq  
TGTGCTAATTCAGGTCCGTCTCCTGCAATCTGTAAATAAACATTTTTCAG Majority  
12110 12120 12130 12140 12150  
11967 TGTGCTAATTCAGGTCCGTCTCCTGCAATCTGTAAATAAACATTTTTCAG 2603\_al2.seq  
11967 TGTGCTAATTCAGGTCCGTCTCCTGCAATCTGTAAATAAACATTTTTCAG nem316\_al2.seq  
GTACTGTGACATCGAAAATGCTTCTAAGAGCAATTCAATGCCTTTTTCTT Majority  
12160 12170 12180 12190 12200  
12017 GTACTGTGACATCGAAAATGCTTCTAAGAGCAATTCAATGCCTTTTTCTT 2603\_al2.seq  
12017 GTACTGTGACATCGAAAATGCTTCTAAGAGCAATTCAATGCCTTTTTCTT nem316\_al2.seq  
TAATAATTCTACCAGCATAAGTGATGAAAATATCATCAGCAGATTTTTC Majority  
12210 12220 12230 12240 12250  
12067 TAATAATTCTACCAGCATAAGTGATGAAAATATCATCAGCAGATTTTTC 2603\_al2.seq  
12067 TAATAATTCTACCAGCATAAGTGATGAAAATATCATCAGCAGATTTTTC nem316\_al2.seq  
AGGTAAGCCGTACCAGCAAAATCAGAGCCTAGACTTTTCAGATACCGAATT Majority  
12260 12270 12280 12290 12300  
12117 AGGTAAGCCGTACCAGCAAAATCAGAGCCTAGACTTTTCAGATACCGAATT 2603\_al2.seq  
12117 AGGTAAGCCGTACCAGCAAAATCAGAGCCTAGACTTTTCAGATACCGAATT nem316\_al2.seq  
ATAAATAACTCCTTTAGCTTCTATATTAATAATGTTTTAACCATTCAACGC Majority  
12310 12320 12330 12340 12350  
12167 AATAAATAACTCCTTTAGCTTCTATATTAATAATGTTTTAACCATTCAACGC 2603\_al2.seq  
12167 AATAAATAACTCCTTTAGCTTCTATATTAATAATGTTTTAACCATTCAACGC nem316\_al2.seq

FIGURE 20R



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TTCTCTTGGATAGCGGATATAAAATCTGGACGATAATGCTTAAACACGCGCT Majority
12360 12370 12380 12390 12400
12217 TTCTCTTGGATACCGCATATAAAATCTGGACGATAATGCTTAAACACGCGCT 2603_al2.seq
12217 TTCTCTTGGATACCGCATATAAAATCTGGACGATAATGCTTAAACACGCGCT nem316_al2.seq

GTGAGAAGATGTTTCATAGATAGCTCCAAAGAAATCTAAAAAACGATTATT Majority
12410 12420 12430 12440 12450
12267 GTGAGAAGATGTTTCATAGATAGCTCCAAAGAAATCTAAAAAACGATTATT 2603_al2.seq
12267 GTGAGAAGATGTTTCATAGATAGCTCCAAAGAAATCTAAAAAACGATTATT nem316_al2.seq

GACAGAAAAATGACTTGACCCATGGTCTAAAAACAATACTAGGTAAATGGT Majority
12460 12470 12480 12490 12500
12317 GACAGAAAAATGACTTGACCCATGGTCTAAAAACAATACTAGGTAAATGGT 2603_al2.seq
12317 GACAGAAAAATGACTTGACCCATGGTCTAAAAACAATACTAGGTAAATGGT nem316_al2.seq

GATTCTTTTGCAAAAGATAGCCCTTCTAGCGTTGTTAACTGAAAACGTGTA Majority
12510 12520 12530 12540 12550
12367 GATTCTTTTGCAAAAGATAGCCCTTCTAGCGTTGTTAACTGAAAACGTGTA 2603_al2.seq
12367 GATTCTTTTGCAAAAGATAGCCCTTCTAGCGTTGTTAACTGAAAACGTGTA nem316_al2.seq

TTACAAATCACAATAATCAATATTTTTCATCTGAAACATATTTTCATCAGCGT Majority
12560 12570 12580 12590 12600
12417 TTACAAATCACAATAATCAATATTTTTCATCTGAAACATATTTTCATCAGCGT 2603_al2.seq
12417 TTACAAATCACAATAATCAATATTTTTCATCTGAAACATATTTTCATCAGCGT nem316_al2.seq

GTTGTATTCTCGATTTTGTGTTAATAATAGGATAGCGCTGCTTGACAATAT Majority
12610 12620 12630 12640 12650
12467 GTTGTATTCTCGATTTTGTGTTAATAATAGGATAGCGCTGCTTGACAATAT 2603_al2.seq
12467 GTTGTATTCTCGATTTTGTGTTAATAATAGGATAGCGCTGCTTGACAATAT nem316_al2.seq

TTTTGGTTCGGTAAACGGTAAATTTTCTACCCTTGTCTTCATCTATAATC Majority
12660 12670 12680 12690 12700
12517 TTTTGGTTCGGTAAACGGTAAATTTTCTACCCTTGTCTTCATCTATAATC 2603_al2.seq
12517 TTTTGGTTCGGTAAACGGTAAATTTTCTACCCTTGTCTTCATCTATAATC nem316_al2.seq

GGTAAATCACCATGATTAGTTGTTACAATAACAACACGGTAGCCACGCTT Majority
12710 12720 12730 12740 12750
12567 GGTAAATCACCATGATTAGTTGTTACAATAACAACACGGTAGCCACGCTT 2603_al2.seq
12567 GGTAAATCACCATGATTAGTTGTTACAATAACAACACGGTAGCCACGCTT nem316_al2.seq

AACCAAATCTGCTGTCATTTTATCTGTATAACGTTCAATACCTCCGAGGA Majority
12760 12770 12780 12790 12800
12617 AACCAAATCTGCTGTCATTTTATCTGTATAACGTTCAATACCTCCGAGGA 2603_al2.seq
12617 AACCAAATCTGCTGTCATTTTATCTGTATAACGTTCAATACCTCCGAGGA nem316_al2.seq

AGGGTAGATAATATCCTGAGAAAACAGCAACTGTTTTTACCTTATTTTCC Majority
12810 12820 12830 12840 12850
12667 AGGGTAGATAATATCCTGAGAAAACAGCAACTGTTTTTACCTTATTTTCC 2603_al2.seq
12667 AGGGTAGATAATATCCTGAGAAAACAGCAACTGTTTTTACCTTATTTTCC nem316_al2.seq

ATATTTATCCACTTTTCATCAATAAGCCATCTTTTAAAGCCTTTAATCATAG Majority
12860 12870 12880 12890 12900
12717 ATATTTATCCACTTTTCATCAATAAGCCATCTTTTAAAGCCTTTAATCATAG 2603_al2.seq
12717 ATATTTATCCACTTTTCATCAATAAGCCATCTTTTAAAGCCTTTAATCATAG nem316_al2.seq

CAACTATTTTTTTGCTCTTTTCTGCTACCAACAACCTCGAACAATA Majority
12910 12920 12930 12940 12950
12767 CAACTATTTTTTTGCTCTTTTCTGCTACCAACAACCTCGAACAATA 2603_al2.seq
12767 CAACTATTTTTTTGCTCTTTTCTGCTACCAACAACCTCGAACAATA nem316_al2.seq

TCATTTTCGCATAAAATACTAAATATTTGTATCGCTTCTTTTACCATATTT Majority
12960 12970 12980 12990 13000
12817 TCATTTTCGCATAAAATACTAAATATTTGTATCGCTTCTTTTACCATATTT 2603_al2.seq
12817 TCATTTTCGCATAAAATACTAAATATTTGTATCGCTTCTTTTACCATATTT nem316_al2.seq

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FIGURE 20S

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TTTTATAAATATAGATCGCATTTGCGTATCATGTAATATTTTCGAAATGGTG Majority  
13010 13020 13030 13040 13050

12867 TTTTATAAATATAGATCGCATTTGCGTATCATGTAATATTTTCGAAATGGTG 2603\_al2.seq  
12867 TTTTATAAATATAGATCGCATTTGCGTATCATGTAATATTTTCGAAATGGTG nem316\_al2.seq

AATGATTCAATACATGAAAAACATGGCCAAATTTTTTAACCTCGTGAAGAG Majority  
13060 13070 13080 13090 13100

12917 AATGATTCAATACATGAAAAACATGGCCAAATTTTTTAACCTCGTGAAGAG 2603\_al2.seq  
12917 AATGATTCAATACATGAAAAACATGGCCAAATTTTTTAACCTCGTGAAGAG nem316\_al2.seq

TGTCCAATTTTCGTGTAACAGACCAATAAAATTAACCTGATAAGTCTTATA Majority  
13110 13120 13130 13140 13150

12967 TGTCCAATTTTCGTGTAACAGACCAATAAAATTAACCTGATAAGTCTTATA 2603\_al2.seq  
12967 TGTCCAATTTTCGTGTAACAGACCAATAAAATTAACCTGATAAGTCTTATA nem316\_al2.seq

TCCCATCTCTGACAGACGATAATTTCATTTTCAGAGTCAACAAAATCAATAA Majority  
13160 13170 13180 13190 13200

13017 TCCCATCTCTGACAGACGATAATTTCATTTTCAGAGTCAACAAAATCAATAA 2603\_al2.seq  
13017 TCCCATCTCTGACAGACGATAATTTCATTTTCAGAGTCAACAAAATCAATAA nem316\_al2.seq

ACATCTCTTCTGCAAAGCCAGATGTTTCTTCGAAAACGCTCGTTTTTCATT Majority  
13210 13220 13230 13240 13250

13067 ACATCTCTTCTGCAAAGCCAGATGTTTCTTCGAAAACGCTCGTTTTTCATT 2603\_al2.seq  
13067 ACATCTCTTCTGCAAAGCCAGATGTTTCTTCGAAAACGCTCGTTTTTCATT nem316\_al2.seq

AAAGCAGCCGAAGTAATACACTCTTCAATTTCTTTATAGTCAAATTCTTG Majority  
13260 13270 13280 13290 13300

13117 AAAGCAGCCGAAGTAATACACTCTTCAATTTCTTTATAGTCAAATTCTTG 2603\_al2.seq  
13117 AAAGCAGCCGAAGTAATACACTCTTCAATTTCTTTATAGTCAAATTCTTG nem316\_al2.seq

CATCACTAAATTTTTCACGGTTTCATATCTTGATACAAACAAGATAACATAC Majority  
13310 13320 13330 13340 13350

13167 CATCACTAAATTTTTCACGGTTTCATATCTTGATACAAACAAGATAACATAC 2603\_al2.seq  
13167 CATCACTAAATTTTTCACGGTTTCATATCTTGATACAAACAAGATAACATAC nem316\_al2.seq

CGACCTTAGGTAAATGAAGGTAATTTTTCATAATTATCTATCAAATCACCT Majority  
13360 13370 13380 13390 13400

13217 CGACCTTAGGTAAATGAAGGTAATTTTTCATAATTATCTATCAAATCACCT 2603\_al2.seq  
13217 CGACCTTAGGTAAATGAAGGTAATTTTTCATAATTATCTATCAAATCACCT nem316\_al2.seq

AGGACAACCCGAATCTTGATCTAAAGTCAAGAACCAATCAAATTCTTGTC Majority  
13410 13420 13430 13440 13450

13267 AGGACAACCCGAATCTTGATCTAAAGTCAAGAACCAATCAAATTCTTGTC 2603\_al2.seq  
13267 AGGACAACCCGAATCTTGATCTAAAGTCAAGAACCAATCAAATTCTTGTC nem316\_al2.seq

TACTGCAAATTTGACCGATACAGTTCAAAGCATATGCAATCCCTTTATTTT Majority  
13460 13470 13480 13490 13500

13317 TACTGCAAATTTGACCGATACAGTTCAAAGCATATGCAATCCCTTTATTTT 2603\_al2.seq  
13317 TACTGCAAATTTGACCGATACAGTTCAAAGCATATGCAATCCCTTTATTTT nem316\_al2.seq

CTGTTAAATAATCAACAGTTAGGTGCCCTCTTTCATTATAATCGGCTACT Majority  
13510 13520 13530 13540 13550

13367 CTGTTAAATAATCAACAGTTAGGTGCCCTCTTTCATTATAATCGGCTACT 2603\_al2.seq  
13367 CTGTTAAATAATCAACAGTTAGGTGCCCTCTTTCATTATAATCGGCTACT nem316\_al2.seq

AATTGAGAAATTTCTTCTTATTTTTTCGAGCCATTATCTACGATATAGAT Majority  
13560 13570 13580 13590 13600

13417 AATTGAGAAATTTCTTCTTATTTTTTCGAGCCATTATCTACGATATAGAT 2603\_al2.seq  
13417 AATTGAGAAATTTCTTCTTATTTTTTCGAGCCATTATCTACGATATAGAT nem316\_al2.seq

GTGGCTTACTTGAGGATAAAATTGCTCGAATGTTCTGATCTAAGCGTTCAA Majority  
13610 13620 13630 13640 13650

13467 GTGGCTTACTTGAGGATAAAATTGCTCGAATGTTCTGATCTAAGCGTTCAA 2603\_al2.seq  
13467 GTGGCTTACTTGAGGATAAAATTGCTCGAATGTTCTGATCTAAGCGTTCAA nem316\_al2.seq

FIGURE 20T



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		T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T T C A T G T T C T A T G C T										Majority
		13660		13670		13680		13690		13700		
13517	T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T T C A T G T T C T A T G C T											2603_al2.seq
13517	T A T T G G G G T T A A A G G T G A C A A T A C C C G C T A A A T A T T T C A T G T T C T A T G C T											nem316_al2.seq
		C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C T G G T G C T T T T G G T T A T A A A A										Majority
		13710		13720		13730		13740		13750		
13567	C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C T G G T G C T T T T G G T T A T A A A A											2603_al2.seq
13567	C T T T T C T A A A A T C T C T A A A T A A C T G A A T G A C T G G T G C T T T T G G T T A T A A A A											nem316_al2.seq
		A C G A T A C C G A C A T A G A T A G T T A C T G C T A C T A A A C T T T G A A T G A C A T A A T T										Majority
		13760		13770		13780		13790		13800		
13617	A C G A T A C C G A C A T A G A T A G T T A C T G C T A C T A A A C T T T G A A T G A C A T A A T T											2603_al2.seq
13617	A C G A T A C C G A C A T A G A T A G T T A C T G C T A C T A A A C T T T G A A T G A C A T A A T T											nem316_al2.seq
		T A C C A A T G A T A C T G A C A T T T G A G T A T T G A T A T A A T A G A G T A C A G C T C C A C										Majority
		13810		13820		13830		13840		13850		
13667	T A C C A A T G A T A C T G A C A T T T G A G T A T T G A T A T A A T A G A G T A C A G C T C C A C											2603_al2.seq
13667	T A C C A A T G A T A C T G A C A T T T G A G T A T T G A T A T A A T A G A G T A C A G C T C C A C											nem316_al2.seq
		T A A G A G T A G C A G C A A T T A A A T A G C G C A G C A T T C C T C T T G T T A A T T C T T T A										Majority
		13860		13870		13880		13890		13900		
13717	T A A G A G T A G C A G C A A T T A A A T A G C G C A G C A T T C C T C T T G T T A A T T C T T T A											2603_al2.seq
13717	T A A G A G T A G C A G C A A T T A A A T A G C G C A G C A T T C C T C T T G T T A A T T C T T T A											nem316_al2.seq
		A A A G T A A A T A C A T C T C T T A A A G A G A T A G C T T G A T A T A G G G A G A C A A T A A A										Majority
		13910		13920		13930		13940		13950		
13767	A A A G T A A A T A C A T C T C T T A A A G A G A T A G C T T G A T A T A G G G A G A C A A T A A A											2603_al2.seq
13767	A A A G T A A A T A C A T C T C T T A A A G A G A T A G C T T G A T A T A G G G A G A C A A T A A A											nem316_al2.seq
		T T C A G T A A T A A C T G T A G A G A T A A T A G C T C C C A T A G C A C C T A A A A T T G G T A										Majority
		13960		13970		13980		13990		14000		
13817	T T C A G T A A T A A C T G T A G A G A T A A T A G C T C C C A T A G C A C C T A A A A T T G G T A											2603_al2.seq
13817	T T C A G T A A T A A C T G T A G A G A T A A T A G C T C C C A T A G C A C C T A A A A T T G G T A											nem316_al2.seq
		T T A A A A G T A T A T T A A A G C A C A A C A T T T G C C A C A A G T C C A A T A A C T G C A G A C										Majority
		14010		14020		14030		14040		14050		
13867	T T A A A A G T A T A T T A A A G C A C A A C A T T T G C C A C A A G T C C A A T A A C T G C A G A C											2603_al2.seq
13867	T T A A A A G T A T A T T A A A G C A C A A C A T T T G C C A C A A G T C C A A T A A C T G C A G A C											nem316_al2.seq
		A T T G T G T A A G C T T T T T G T A C G T C T T G A A G C C A G T A G A T A C T G T G T C C C T A A										Majority
		14060		14070		14080		14090		14100		
13917	A T T G T G T A A G C T T T T T G T A C G T C T T G A A G C C A G T A G A T A C T G T G T C C C T A A											2603_al2.seq
13917	A T T G T G T A A G C T T T T T G T A C G T C T T G A A G C C A G T A G A T A C T G T G T C C C T A A											nem316_al2.seq
		A G C G T T A C C A T A A G A A A T G C A A A T G A T C A T C A A A										Majority
		14110		14120		14130						
13967	A G C G T T A C C A T A A G A A A T G C A A A T G A T C A T C A A A											2603_al2.seq
13967	A G C G T T A C C A T A A G A A A T G C A A A T G A T C A T C A A A											nem316_al2.seq

Decoration 'Decoration #1': Shade (with solid black) residues that differ from the Consensus.

FIGURE 20U

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TCCACATCGGTCCCAATTAACATATGACGTGGCGCATCACCAGTAATTTCGG Majority
10 20 30 40 50
1 TCCACATCGGTCCCAATTAACATATGACGTGGCGCATCACCAGTAATTTCGG coh1_al2.seq
1 TCCACATCGGTCCCAATTAACATATGACGTGGCGCATCACCAGTAATTTCGG a909_al2.seq

TGAATAACAATATGTTTTTGGGAATAATCTCCAGTTGGTTCACAAATAATCGA Majority
60 70 80 90 100
51 TGAATAACAATATGTTTTTGGGAATAATCTCCAGTTGGTTCACAAATAATCGA coh1_al2.seq
51 TGAATAACAATATGTTTTTGGGAATAATCTCCAGTTGGTTCACAAATAATCGA a909_al2.seq

AATATAGTCTTCTTGACTTAACAAACGTAAACGACCTTCATGGTAATCTC Majority
110 120 130 140 150
101 AATATAGTCTTCTTGACTTAACAAACGTAAACGACCTTCATGGTAATCTC coh1_al2.seq
101 AATATAGTCTTCTTGACTTAACAAACGTAAACGACCTTCATGGTAATCTC a909_al2.seq

TCTGCATTCTTGTATTAGTTCATAAGATGCAGAAGGTGTAATTTTATACCC Majority
160 170 180 190 200
151 TCTGCATTCTTGTATTAGTTCATAAGATGCAGAAGGTGTAATTTTATACCC coh1_al2.seq
151 TCTGCATTCTTGTATTAGTTCATAAGATGCAGAAGGTGTAATTTTATACCC a909_al2.seq

TGAATATCATTATCCGTAACACATCGACGAACATTTTCCACCATCATATC Majority
210 220 230 240 250
201 TGAATATCATTATCCGTAACACATCGACGAACATTTTCCACCATCATATC coh1_al2.seq
201 TGAATATCATTATCCGTAACACATCGACGAACATTTTCCACCATCATATC a909_al2.seq

ATGTGTCTCCCTGGGAGACCATTATTAGGTGAGAAACGATTTCTACTT Majority
260 270 280 290 300
251 ATGTGTCTCCCTGGGAGACCATTATTAGGTGAGAAACGATTTCTACTT coh1_al2.seq
251 ATGTGTCTCCCTGGGAGACCATTATTAGGTGAGAAACGATTTCTACTT a909_al2.seq

TAGGAGCTAATTCTCGTATTCTCTTAACAGTTTTTTTGTAAAGGTCATAT Majority
310 320 330 340 350
301 TAGGAGCTAATTCTCGTATTCTCTTAACAGTTTTTTTGTAAAGGTCATAT coh1_al2.seq
301 TAGGAGCTAATTCTCGTATTCTCTTAACAGTTTTTTTGTAAAGGTCATAT a909_al2.seq

GAATGTCTCTATTTTATTAATGCAGAAGTTGCTTCATAAGTTGTCTGAAG Majority
360 370 380 390 400
351 GAATGTCTCTATTTTATTAATGCAGAAGTTGCTTCATAAGTTGTCTGAAG coh1_al2.seq
351 GAATGTCTCTATTTTATTAATGCAGAAGTTGCTTCATAAGTTGTCTGAAG a909_al2.seq

GCCTAATTCTAAAGTCACATGCATTCTTTTCAGAAAGTTTCAGCCGAGATAGT Majority
410 420 430 440 450
401 GCCTAATTCTAAAGTCACATGCATTCTTTTCAGAAAGTTTCAGCCGAGATAGT coh1_al2.seq
401 GCCTAATTCTAAAGTCACATGCATTCTTTTCAGAAAGTTTCAGCCGAGATAGT a909_al2.seq

ATATAGTTTTTCATCAGGTAAGCAATCCGGCCTTGTTCGGATGTTGATCCCC Majority
460 470 480 490 500
451 ATATAGTTTTTCATCAGGTAAGCAATCCGGCCTTGTTCGGATGTTGATCCCC coh1_al2.seq
451 ATATAGTTTTTCATCAGGTAAGCAATCCGGCCTTGTTCGGATGTTGATCCCC a909_al2.seq

ATAACTCCTGGCTCATTAAATAGCCTGTTCGTAACGCTCTTTAATTATCTC Majority
510 520 530 540 550
501 ATAACTCCTGGCTCATTAAATAGCCTGTTCGTAACGCTCTTTAATTATCTC coh1_al2.seq
501 ATAACTCCTGGCTCATTAAATAGCCTGTTCGTAACGCTCTTTAATTATCTC a909_al2.seq

TAACCTTAGCATGGGTATTGGTAAATTTTGAATAAGACTAAGTATTTAT Majority
560 570 580 590 600
551 TAACCTTAGCATGGGTATTGGTAAATTTTGAATAAGACTAAGTATTTAT coh1_al2.seq
551 TAACCTTAGCATGGGTATTGGTAAATTTTGAATAAGACTAAGTATTTAT a909_al2.seq

TAACCTCAGGCCACTTTCTATGCATGAAATCAATTTCTTTATAGAATTGT Majority
610 620 630 640 650
601 TAACCTCAGGCCACTTTCTATGCATGAAATCAATTTCTTTATAGAATTGT coh1_al2.seq
601 TAACCTCAGGCCACTTTCTATGCATGAAATCAATTTCTTTATAGAATTGT a909_al2.seq
```

Figure 21



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TCACGAATAGGAGCTTCTGAGCAACTATAGCATCCCTGAAACCAGAAAC Majority  
660 670 680 690 700

651 TCACGAATAGGAGCTTCTGAGCAACTATAGCATCCCTGAAACCAGAAAC coh1\_al2.seq  
651 TCACGAATAGGAGCTTCTGAGCAACTATAGCATCCCTGAAACCAGAAAC a909\_al2.seq

TGTGCAAAAAGTGCAACCCTCCTCTAGCAACTGTTCCATCTCTGTTAGGAC Majority  
710 720 730 740 750

701 TGTGCAAAAAGTGCAACCCTCCTCTAGCAACTGTTCCATCTCTGTTAGGAC coh1\_al2.seq  
701 TGTGCAAAAAGTGCAACCCTCCTCTAGCAACTGTTCCATCTCTGTTAGGAC a909\_al2.seq

AGTCAAAACCAGCATCTATAGGTAATTTAAATATTTTTTCTCCAAAGAGT Majority  
760 770 780 790 800

751 AGTCAAAACCAGCATCTATAGGTAATTTAAATATTTTTTCTCCAAAGAGT coh1\_al2.seq  
751 AGTCAAAACCAGCATCTATAGGTAATTTAAATATTTTTTCTCCAAAGAGT a909\_al2.seq

TCTCGATAATAATCATTAAATCGCACGATAACGTTTTTTCATAGGATAATT Majority  
810 820 830 840 850

801 TCTCGATAATAATCATTAAATCGCACGATAACGTTTTTTCATAGGATAATT coh1\_al2.seq  
801 TCTCGATAATAATCATTAAATCGCACGATAACGTTTTTTCATAGGATAATT a909\_al2.seq

GTATCACAATTTTAACTAAAATAACCTCACTACTACAATAAAACTAAAAA Majority  
860 870 880 890 900

851 GTATCACAATTTTAACTAAAATAACCTCACTACTACAATAAAACTAAAAA coh1\_al2.seq  
851 GTATCACAATTTTAACTAAAATAACCTCACTACTACAATAAAACTAAAAA a909\_al2.seq

AGATTGGAACGTCAGTTAGTTCCAATCTTTTATTTACTTCACTTTCTTTA Majority  
910 920 930 940 950

901 AGATTGGAACGTCAGTTAGTTCCAATCTTTTATTTACTTCACTTTCTTTA coh1\_al2.seq  
901 AGATTGGAACGTCAGTTAGTTCCAATCTTTTATTTACTTCACTTTCTTTA a909\_al2.seq

ACCAATCCTTGGCTAAAAAGATATACGCAGTTAGATTCAAAATACCATAA Majority  
960 970 980 990 1000

951 ACCAATCCTTGGCTAAAAAGATATACGCAGTTAGATTCAAAATACCATAA coh1\_al2.seq  
951 ACCAATCCTTGGCTAAAAAGATATACGCAGTTAGATTCAAAATACCATAA a909\_al2.seq

GCAAGTATAAAACCAGCTAAAACATCTGTGCGGAAAATGAACCCCTAGGTA Majority  
1010 1020 1030 1040 1050

1001 GCAAGTATAAAACCAGCTAAAACATCTGTGCGGAAAATGAACCCCTAGGTA coh1\_al2.seq  
1001 GCAAGTATAAAACCAGCTAAAACATCTGTGCGGAAAATGAACCCCTAGGTA a909\_al2.seq

AATACGAGATAACCCAATTAAAAAAATGAGCAAACCCAATGTACCTTGGC Majority  
1060 1070 1080 1090 1100

1051 AATACGAGATAACCCAATTAAAAAAATGAGCAAACCCAATGTACCTTGGC coh1\_al2.seq  
1051 AATACGAGATAACCCAATTAAAAAAATGAGCAAACCCAATGTACCTTGGC a909\_al2.seq

ACAACAGTTTCCATATACTCTTAGGGCATATAGTACTGCAATAAAATAATA Majority  
1110 1120 1130 1140 1150

1101 ACAACAGTTTCCATATACTCTTAGGGCATATAGTACTGCAATAAAATAATA coh1\_al2.seq  
1101 ACAACAGTTTCCATATACTCTTAGGGCATATAGTACTGCAATAAAATAATA a909\_al2.seq

CTACTCCCAAATATCATAAATGTTCCCATCGAGTCCCCACTGGGAAACGA Majority  
1160 1170 1180 1190 1200

1151 CTACTCCCAAATATCATAAATGTTCCCATCGAGTCCCCACTGGGAAACGA coh1\_al2.seq  
1151 CTACTCCCAAATATCATAAATGTTCCCATCGAGTCCCCACTGGGAAACGA a909\_al2.seq

ATAGCCACCTGCAAATACTAAATGGGTTAAAGTTGGTCTCACTCTTTGAA Majority  
1210 1220 1230 1240 1250

1201 ATAGCCACCTGCAAATACTAAATGGGTTAAAGTTGGTCTCACTCTTTGAA coh1\_al2.seq  
1201 ATAGCCACCTGCAAATACTAAATGGGTTAAAGTTGGTCTCACTCTTTGAA a909\_al2.seq

AAATAAGTTTTTAAAGAAAGTATACATATACCAGAGATAATAGCATTTACT Majority  
1260 1270 1280 1290 1300

1251 AAATAAGTTTTTAAAGAAAGTATACATATACCAGAGATAATAGCATTTACT coh1\_al2.seq  
1251 AAATAAGTTTTTAAAGAAAGTATACATATACCAGAGATAATAGCATTTACT a909\_al2.seq

FIGURE 21A

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	G C G A T A A A A T C T A G C T T G A G G A T A C C A C T T C T T A A G G T A A C A G A A A G T G A C	Majority
	1310 1320 1330 1340 1350	
1301	G C G A T A A A A T C T A G C T T G A G G A T A C C A C T T C T T A A G G T A A C A G A A A G T G A C	coh1_al2.seq
1301	G C G A T A A A A T C T A G C T T G A G G A T A C C A C T T C T T A A G G T A A C A G A A A G T G A C	a909_al2.seq
	G C T C A T A A A T C G C A A T A G C T A T C T G G C T T A C A G T A T T A C C A A C C A C A G T G A	Majority
	1360 1370 1380 1390 1400	
1351	G C T C A T A A A T C G C A A T A G C T A T C T G G C T T A C A G T A T T A C C A A C C A C A G T G A	coh1_al2.seq
1351	G C T C A T A A A T C G C A A T A G C T A T C T G G C T T A C A G T A T T A C C A A C C A C A G T G A	a909_al2.seq
	T T A A C T T G A A A A A T C T T G T A G A A A G A T T T G G C A A C T G T C C T C T A A C A C T T	Majority
	1410 1420 1430 1440 1450	
1401	T T A A C T T A A A A A A T C T T G T A G A A A G A T T T G G C A A C T G T C C T C T A A C A C T T	coh1_al2.seq
1401	T T A A C T T G A A A A A T C T T G T A G A A A G A T T T G G C A A C T G T C C T C T A A C A C T T	a909_al2.seq
	T C T T G A A T A G T T T G G T C A A A T G C G A T T A C A G T G T C G G G C C A A T A T T T G A T	Majority
	1460 1470 1480 1490 1500	
1451	T C T T G A A T A G T T T G G T C A A A T G C G A T T A C A G T G T C G G G C C A A T A T T T G A T	coh1_al2.seq
1451	T C T T G A A T A G T T T G G T C A A A T G C G A T T A C A G T G T C G G G C C A A T A T T T G A T	a909_al2.seq
	G A C C A A T C C T A A A C T G A A A A A T A A G A T A A T A G C A A T A A A T G C T T G A A T A A	Majority
	1510 1520 1530 1540 1550	
1501	G A C C A A T C C T A A A C T G A A A A A T A A G A T A A T A G C A A T A A A T G C T T G A A T A A	coh1_al2.seq
1501	G A C C A A T C C T A A A C T G A A A A A T A A G A T A A T A G C A A T A A A T G C T T G A A T A A	a909_al2.seq
	G T T T A C T A T T T T G A C G A G A T A A G A T T A G T C T T T T T A T A T C T T T C T A A T A T	Majority
	1560 1570 1580 1590 1600	
1551	G T T T A C T A T T T T G A C G A G A T A A G A T T A G T C T T T T T A T A T C T T T C T A A T A T	coh1_al2.seq
1551	G T T T A C T A T T T T G A C G A G A T A A G A T T A G T C T T T T T A T A T C T T T C T A A T A T	a909_al2.seq
	T G G C A A A C A A G C C A C G T A A G T T A G A T A G A A A C A A T C G A A A T T A A A A T T C	Majority
	1610 1620 1630 1640 1650	
1601	T G G C A A A C A A G C C A C G T A A G T T A G A T A G A A A C A A T C G A A A T T A A A A T T C	coh1_al2.seq
1601	T G G C A A A C A A G C C A C G T A A G T T A G A T A G A A A C A A T C G A A A T T A A A A T T C	a909_al2.seq
	C C T C A A C G A T A T T A A A T G G A A T A A C C A T T G T T A A A A G G T A A T T G C C T A C A	Majority
	1660 1670 1680 1690 1700	
1651	C C T C A A C G A T A T T A A A T G G A A T A A C C A T T G T T A A A A G G T A A T T G C C T A C A	coh1_al2.seq
1651	C C T C A A C G A T A T T A A A T G G A A T A A C C A T T G T T A A A A G G T A A T T G C C T A C A	a909_al2.seq
	C C A A T A A A T G T T C T G A T A T C A A A G T T A G C A A A T A T A G C A T A C A A A G G A A T	Majority
	1710 1720 1730 1740 1750	
1701	C C A A T A A A T G T T C T G A T A T C A A A G T T A G C A A A T A T A G C A T A C A A A G G A A T	coh1_al2.seq
1701	C C A A T A A A T G T T C T G A T A T C A A A G T T A G C A A A T A T A G C A T A C A A A G G A A T	a909_al2.seq
	C G C A A A G A C A T A G T T G A G A G C T A C C A T A G A T A C A G T C A A G C T A A C T G T A C	Majority
	1760 1770 1780 1790 1800	
1751	C G C A A A G A C A T A G T T G A G A G C T A C C A T A G A T A C A G T C A A G C T A A C T G T A C	coh1_al2.seq
1751	C G C A A A G A C A T A G T T G A G A G C T A C C A T A G A T A C A G T C A A G C T A A C T G T A C	a909_al2.seq
	G A A A T A A A C T A G C T T T A A T A A A A T C T T T T G C A C T C T C T A T T T T T C C A G	Majority
	1810 1820 1830 1840 1850	
1801	G A A A T A A A C T A G C T T T A A T A A A A T C T T T T G C A C T C T C T A T T T T T C C A G	coh1_al2.seq
1801	G A A A T A A A C T A G C T T T A A T A A A A T C T T T T G C A C T C T C T A T T T T T C C A G	a909_al2.seq
	A A A A T A G C G A A A C T T G C T A A A A A T A G A G C T A G A G C A A C C A T A T T C A T C G G	Majority
	1860 1870 1880 1890 1900	
1851	A A A A T A G C G A A A C T T G C T A A A A A T A G A G C T A G A G C A A C C A T A T T C A T C G G	coh1_al2.seq
1851	A A A A T A G C G A A A C T T G C T A A A A A T A G A G C T A G A G C A A C C A T A T T C A T C G G	a909_al2.seq
	T A A A C C G A T A A A G G T T T C T G G A C C A C G A T T A G C A A G T A T A A C T T T T A A A A	Majority
	1910 1920 1930 1940 1950	
1901	T A A A C C G A T A A A G G T T T C T G G A C C A C G A T T A G C A A G T A T A A C T T T T A A A A	coh1_al2.seq
1901	T A A A C C G A T A A A G G T T T C T G G A C C A C G A T T A G C A A G T A T A A C T T T T A A A A	a909_al2.seq

FIGURE 21B



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	GTGATCTTAATAAAGAGTACACCAATAAAGTTGATTTCAAATCAAATAAAATA	Majority
	1960 1970 1980 1990 2000	
1951	GTGATCTTAATAAAGAGTACACCAATAAAGTTGATTTCAAATCAAATAAAATA	coh1_al2.seq
1951	GTGATCTTAATAAAGAGTACACCAATAAAGTTGATTTCAAATCAAATAAAATA	a909_al2.seq
	AAAGCAACTAACATCGGAAGGATTGAAAAATCAACCTTTAAAAATTCTGC	Majority
	2010 2020 2030 2040 2050	
2001	AAAGCAACTAACATCGGAAGGATTGAAAAATCAACCTTTAAAAATTCTGC	coh1_al2.seq
2001	AAAGCAACTAACATCGGAAGGATTGAAAAATCAACCTTTAAAAATTCTGC	a909_al2.seq
	TCCTGGTATTAAATGGAAATGAAACCATCATCAATACAAAAGATAAGGCCAG	Majority
	2060 2070 2080 2090 2100	
2051	TCCTGGTATTAAATGGAAATGAAACCATCATCAATACAAAAGATAAGGCCAG	coh1_al2.seq
2051	TCCTGGTATTAAATGGAAATGAAACCATCATCAATACAAAAGATAAGGCCAG	a909_al2.seq
	AAAGAATGGCGATTGTACCAATTTTACGTGTATTGTTCATAAAAAAATTTC	Majority
	2110 2120 2130 2140 2150	
2101	AAAGAATGGCGATTGTACCAATTTTACGTGTATTGTTCATAAAAAAATTTC	coh1_al2.seq
2101	AAAGAATGGCGATTGTACCAATTTTACGTGTATTGTTCATAAAAAAATTTC	a909_al2.seq
	CTCCAATTTAAATAAAATTGAAAGAAGCTCCAAAGGTAAGCGTAGGTACGC	Majority
	2160 2170 2180 2190 2200	
2151	CTCCAATTTAAATAAAATTGAAAGAAGCTCCAAAGGTAAGCGTAGGTACGC	coh1_al2.seq
2151	CTCCAATTTAAATAAAATTGAAAGAAGCTCCAAAGGTAAGCGTAGGTACGC	a909_al2.seq
	GAAAAAAACCTTTGTCTTCTCCCATCCAGACTTTACTGTCTGGTTGTGGA	Majority
	2210 2220 2230 2240 2250	
2201	GAAAAAAACCTTTGTCTTCTCCCATCCAGACTTTACTGTCTGGTTGTGGA	coh1_al2.seq
2201	GAAAAAAACCTTTGTCTTCTCCCATCCAGACTTTACTGTCTGGTTGTGGA	a909_al2.seq
	TCTCACCACATCAGCTTTTCGCTCGCGGACTGATGCTTCACAACTGACAAA	Majority
	2260 2270 2280 2290 2300	
2251	TCTCACCACATCAGCTTTTCGCTCGCGGACTGATGCTTCACAACTGACAAA	coh1_al2.seq
2251	TCTCACCACATCAGCTTTTCGCTCGCGGACTGATGCTTCACAACTGACAAA	a909_al2.seq
	TAAAGTTGGAAGCGATTACCGCCGGTCCGGGAATTACACCCTGCCCTGAAGA	Majority
	2310 2320 2330 2340 2350	
2301	TAAAGTTGGAAGCGATTACCGCCGGTCCGGGAATTACACCCTGCCCTGAAGA	coh1_al2.seq
2301	TAAAGTTGGAAGCGATTACCGCCGGTCCGGGAATTACACCCTGCCCTGAAGA	a909_al2.seq
	CACCTATAGCATAACAAAAAAACTTTGCAATTGCAAGTTTTTTAATCACT	Majority
	2360 2370 2380 2390 2400	
2351	CACCTATAGCATAACAAAAAAACTTTGCAATTGCAAGTTTTTTAATCACT	coh1_al2.seq
2351	CACCTATAGCATAACAAAAAAACTTTGCAATTGCAAGTTTTTTAATCACT	a909_al2.seq
	AATTAGTAGTAGATTGTATAATATTAATTTTTTAACATCAATTAATTGACA	Majority
	2410 2420 2430 2440 2450	
2401	AATTAGTAGTAGATTGTATAATATTAATTTTTTAACATCAATTAATTGACA	coh1_al2.seq
2401	AATTAGTAGTAGATTGTATAATATTAATTTTTTAACATCAATTAATTGACA	a909_al2.seq
	GCGCACTAATACTCTAGCTACTCCTGCCCTTTGTACAAAGTAAACAAGCTTA	Majority
	2460 2470 2480 2490 2500	
2451	GCGCACTAATACTCTAGCTACTCCTGCCCTTTGTACAAAGTAAACAAGCTTA	coh1_al2.seq
2451	GCGCACTAATACTCTAGCTACTCCTGCCCTTTGTACAAAGTAAACAAGCTTA	a909_al2.seq
	AGTCCCAATCATTGTCTGATGTGGCAGTTTTTATAAACTTTTTCAATCGCT	Majority
	2510 2520 2530 2540 2550	
2501	AGTCCCAATCATTGTCTGATGTGGCAGTTTTTATAAACTTTTTCAATCGCT	coh1_al2.seq
2501	AGTCCCAATCATTGTCTGATGTGGCAGTTTTTATAAACTTTTTCAATCGCT	a909_al2.seq
	GTTGGTTCAATAAATTTCTCTATTACTGATTTTGTAGTGATAGATTTGCCC	Majority
	2560 2570 2580 2590 2600	
2551	GTTGGTTCAATAAATTTCTCTATTACTGATTTTGTAGTGATAGATTTGCCC	coh1_al2.seq
2551	GTTGGTTCAATAAATTTCTCTATTACTGATTTTGTAGTGATAGATTTGCCC	a909_al2.seq

FIGURE 21C

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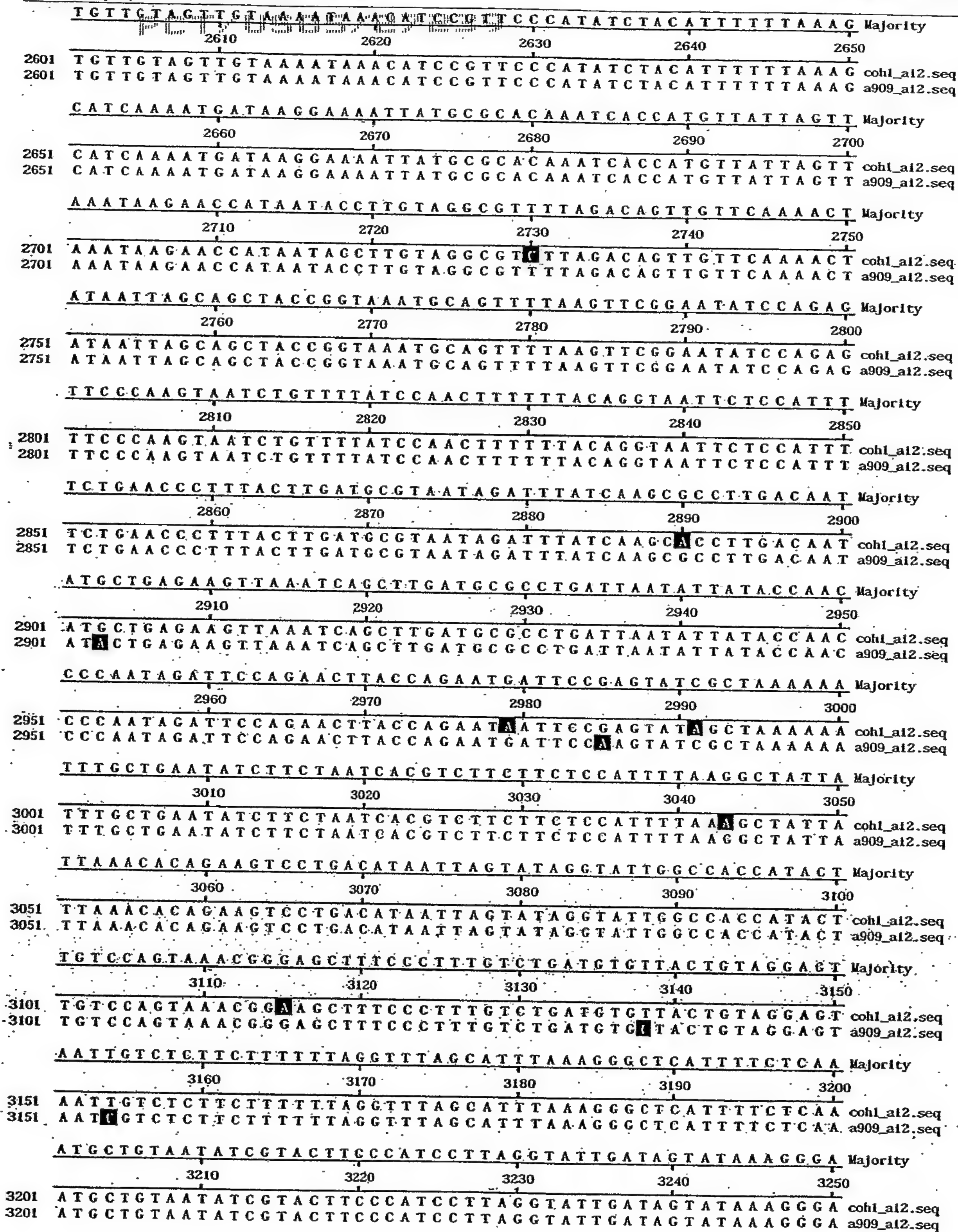


FIGURE 21D



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	G A C A T T A G T T C A T A A C C T T G A G C T G T T T A G T C T G A A T A A A T A G A T A A A T	Majority
	3260 3270 3280 3290 3300	
3251	G A C A T T A G T T C A T A A C C T T G A G C T G T T T T A G T C T G A A T A A A T A G A T A A A T	cohl_al2.seq
3251	G A C A T T A G T T C A T A A C C T T G A G C T G T T T T A G T C T G A A T A A A T A G A T A A A T	a909_al2.seq
	C C C T T G A G G A A G A T T G T T C G C A A C A A T A C C T T C A G C C G G T A A A T T A T C A A	Majority
	3310 3320 3330 3340 3350	
3301	C C C T T G A G G A A G A T T G T T C G C A A C A A T A C C T T C A G C C G G T A A A T T A T C A A	cohl_al2.seq
3301	C C C T T G A G G A A G A T T G T T C G C A A C A A T A C C T T C A G C C G G T A A A T T A T C A A	a909_al2.seq
	A C G T T T G T A A A G G T T G A G T T T T A T G A A C A G C T T T T G T T A G T A G A T T G A C G	Majority
	3360 3370 3380 3390 3400	
3351	A C G T T T G T A A A G G T T G A G T T T T A T G A A C A G C T T T T G T T A G T A G A T T G A C G	cohl_al2.seq
3351	A C G T T T G T A A A G G T T G A T T T T T A T G A A C A G C T T T T G T T A G T A G A T T G A C A	a909_al2.seq
	T A T T T G G C T T G G T T A C T A T C A A G G T T T A C T T G T G T T A G A T C A T C G T C T T T	Majority
	3410 3420 3430 3440 3450	
3401	T A T T T G G C T T G G T T A C T A T C A A G G T T T A C T T G T G T T A G A T C A T C G T C T T T	cohl_al2.seq
3401	T A T T T G G C T T G G T T A C T A T C A A G G T T T A C T T G T G T T A A T C A T C G T C T T T	a909_al2.seq
	T A T T C C A A T A C C T T G A A A T G G G G T A G T T A G A G T A A A A A C T T G G T T A C C A T	Majority
	3460 3470 3480 3490 3500	
3451	T A T T C C A A T A C C T T G A A A T G G G G T A G T T A G A G T A A A A A C T T G A T T A C C A T	cohl_al2.seq
3451	T A T T C C A A T A C C T T G A A A T G G G G T A G T T A G A G T A A A A A C T T G G T T A C C A T	a909_al2.seq
	C A A C A T C T T T A G C T T G T G C T A C T T G G T A A A C A A G T A A A T T A C C G C C A G C G	Majority
	3510 3520 3530 3540 3550	
3501	C A A C A T C T T T A G C T T G T G C T A C T T G G T A A A C A A G T A A A T T A C C G C C A G C G	cohl_al2.seq
3501	C A A C A T C T T T A G C T T G T G C T A C T T G G T A A A C A A G T A A A T T A C C G C C A G C G	a909_al2.seq
	A T A C C T T G A T T A T T A T A C T T A T T T T G T A T A G T A A T A G A A C C C G T T T T C A T	Majority
	3560 3570 3580 3590 3600	
3551	A T A C C T T G A T T A T T A T A C T T A T T T T G T A T A G T A A T A G A A C C C G T T T T C A T	cohl_al2.seq
3551	A T A C C T T G A T T A T T A T A C T T A T T T T G T A T A G T A A T A G A A C C C G T T T T C A T	a909_al2.seq
	C T G A T C A T T G G T A T C A G C A G A C A C A A G T T G A G T A C T T A G A C T A A A T A A T A	Majority
	3610 3620 3630 3640 3650	
3601	C T G A T C A T T G G T A T C A G C A G A C A C A A G T T G A G T A C T T A G A C T A A A T A A T A	cohl_al2.seq
3601	C T G A T C A T T G G T A T C A G C A G A C A C A A G T T G A G T A C T T A G A C T A A A T A A T A	a909_al2.seq
	A G A G A A G A G T T A T C T T T A G G A T C T T T T T A T A A A T C A T T G T T C T C T T C C T T	Majority
	3660 3670 3680 3690 3700	
3651	A G A G A A G A G T T A T C T T T A G G A T C T T T T T A T A A A T C A T T G T T C T C T T C C T T	cohl_al2.seq
3651	A G A G A A G A G T T A T A T T T A G G A T C T T T T T A T A A A T C A T T G T T C T C T T C C T T	a909_al2.seq
	T C T C A T T G C T T G T T T T A A A A T T T T C T T A C G T T G A C G T G C T C T C C T A G T T A	Majority
	3710 3720 3730 3740 3750	
3701	T C T C A T T G C T T G T T T T A A A A T T T T C T T A C G T T G A C G T G C T C T C C T A G T T A	cohl_al2.seq
3701	T C T C A T T G C T T G T T T T A A A A T T T T C T T A C G T T G A C G T G C T C T C C T A G T T A	a909_al2.seq
	C T T C T A A A G A G A T T A A A A G T A A A A T C A A A G T A A G G A A A A T A G C G A T A A A T	Majority
	3760 3770 3780 3790 3800	
3751	C T T C T A A A G A G A T T A A A A G T A A A A T C A A A G T A A G G A A A A T A G C G A T A A A T	cohl_al2.seq
3751	C T T C T A A A A G A T T A A A A G T A A A A T C A A A G T A A G G A A A A T A G C G A T A A A T	a909_al2.seq
	G G T G C G A T A T A A A T A G G C T C T A T T T G T A T T G C C T C T G C T A C T A C C A A A G C	Majority
	3810 3820 3830 3840 3850	
3801	G G T G C G A T A T A A A T A G G C T C T A T T T G T A T T G C C T C T G C T A C T A C C A A A G C	cohl_al2.seq
3801	G G T G C G A T A T A A A T A G G C T C T A T T T G T A T T G C C T C T G C T A C T A C C A A A G C	a909_al2.seq
	G T T A C C A T T A T C G T T T G G T A C A C G A T G T C C T C T C A C T A G T A A C C G A T G G G	Majority
	3860 3870 3880 3890 3900	
3851	G T T A C C A T T A T C G T T T G G T A C A C G A T G T C C T C T C A C T A G T A A C C G A T G G G	cohl_al2.seq
3851	G T T A C C A T T A T C G T T T G G T A C A C G A T G T C C T C T C A C T A G T A A C C G A T G G G	a909_al2.seq

FIGURE 21E

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	T A T T A A C G C C A T A T G G T G T A C A C G T C A C C A A A G T T T G G T A G T C T T T A C C T	Majority
	3910 3920 3930 3940 3950	
3901	T A T T A A C G C C A T A T G G T G T A C A C G T C A C C A A A G T T T G G T A G T C T T T A C C T	coh1_al2.seq
3901	T A T T A A C A C C A T A T G G T G T A C A C G T C A C C A A A G T T T G G T A G T C T T T A C C T	a909_al2.seq
	T T A A C A A T T T T G T A A A T C C C T C A A A T C A T C C G G T T T A A C T G T T C T G A T T T G	Majority
	3960 3970 3980 3990 4000	
3951	T T A A C A A T T T T G T A A A T C C C T C A A A T C A T C C G G T T T A A C T G T T C T G A T T T G	coh1_al2.seq
3951	T T A A C A A T T T T G T A A A T C C C T C A A A T C A T C C G G T T T A A C T G T T C T G A T T T G	a909_al2.seq
	A T C C A C T T G A T A A G T A T A T G T T T C A T T T A A G A T A C T G A C T G T C C A G T G G T	Majority
	4010 4020 4030 4040 4050	
4001	A T C C A C T T G A T A A G T A T A T G T T T C A T T T A A G A T A C T G A C T G T C C A G T G G T	coh1_al2.seq
4001	A T C C A C T T G A T A A G T A T A T G T T T C A T T T A A G A T A C T G A C T G T C C A G T G G T	a909_al2.seq
	C T C C A G C T T T T A A C T T A T C C A A A T C A G A A A A A G C C T T G A A G A G G G T A A A	Majority
	4060 4070 4080 4090 4100	
4051	C T C C A A C T T T T A A C T T A T C C A A A T C A G A A A A A A G C C T T G A A G A G G G T A A A	coh1_al2.seq
4051	C T C C A G C T T T T A A C T T A T C C A A A T C A G A A A A A A G C C T T G A A G A G G G T A A A	a909_al2.seq
	C C T C T A T G T C C T G A T A A A A T A G A A T G A G T T G A G T C T C C T C C A A T T G G A A G	Majority
	4110 4120 4130 4140 4150	
4101	C C T C T A T G T C C T G A T A A A A T A G A A T G A G T T G A G T C T C C T C C A A T T G G A A G	coh1_al2.seq
4101	C C T C T A T G T C C T G A T A A A A T A G A A T G A G T T G A G T C T C C T C C A A T T G G A A G	a909_al2.seq
	A C T A C T T C C T T C T A A A T G A C C A A T A G A A G T T T G A A G C A C T T T T T C A C T T G	Majority
	4160 4170 4180 4190 4200	
4151	A C T A C T T C C T T C T A A A T G A C C A A T A G A A G T T T G A A G C A C T T T T T C A C T T G	coh1_al2.seq
4151	A C T A C T T C C T T C T A A A T G A C C A A T A G A A G T T T G A A G C A C T T T T T C A C T T G	a909_al2.seq
	T A C C A T G A T A A A G T G G T A A T T T T A T G T T T A T C T T T G G A A T T G A A A T A T A A	Majority
	4210 4220 4230 4240 4250	
4201	T A C C A T G A T A A A G T G G T A A T T T T A T G T T T A T C T T T G G A A T T G A A A T A T A A	coh1_al2.seq
4201	T A C C A T G A T A A A G T G G T A A T T T T A T A T T T A T C T T T G G A A T T G A A A T A T A A	a909_al2.seq
	C C C A T A T T A C C C G T T T T A T C G A T A G C C A G T T G T G A A T T A T A A T C C A A A C G	Majority
	4260 4270 4280 4290 4300	
4251	C C C A T A T T A C C C G T T T T A T C G A T A G C C A G T T G T G A A T T A T A A T C C A A A C G	coh1_al2.seq
4251	C C C A T A T T A C C C G T T T T A T C G A T A G C C A G T T G T G A A T T A T A A T C C A A A C G	a909_al2.seq
	E T C T E G G T T A G T C A T G T G C C A C T T C A T T C C T G A A G T T T T A A A T T G C T T A T	Majority
	4310 4320 4330 4340 4350	
4301	C T C T T G G T T A G T C A T G T G C C A C T T C A T T C C T G A A G T T T T A A A T T G C T T A T	coh1_al2.seq
4301	C T C T T G G T T A G T C A T G T G C C A C T T C A T T C C T G A A G T T T T A A A T T G C T T A T	a909_al2.seq
	T A T A T T C T T T G G C T C G G T T A A T A A T T T T T T A T A G T C G T T T T C A T C C A T A	Majority
	4360 4370 4380 4390 4400	
4351	T A T A T T C T T T G G C T C G G T T A A T A A T T T T T T A T A T C G T T T T C A T C C A T A	coh1_al2.seq
4351	T A T A T T C T T T G G C T C G G T T A A T A A T T T T T T A T A T C G T T T T C A T C C A T A	a909_al2.seq
	T G C G T T A C G C G G T C T T G G T A A T C C A T A A T C G C T C G A G A T T G G T G A A A T G A	Majority
	4410 4420 4430 4440 4450	
4401	T G C G T T A C G C G G T C T T G G T A A T C C A T A A T C G C T C G A G A T T G G T G A A A T G A	coh1_al2.seq
4401	T G C G T T A C G C G G T C T T G G T A A T C C A T A A T C G C T C G A G A T T G G T G A A A T G A	a909_al2.seq
	A T T C C A A T A A T T A G C A A G T G A A G G A T A A G C C A T T A A G C C T A C C C C C A C T G	Majority
	4460 4470 4480 4490 4500	
4451	A T T C C A A T A A T T A G C A A G T G A A G G A T A A G C C A T T A A G C C T A C C C C C A C T G	coh1_al2.seq
4451	A T T C C A A T A A T T A G C A A G T G A A G G A T A A G C C A T T A A G C C T A C C C C C A C T G	a909_al2.seq
	C A A T T A T A G T G A C A A G C A A A A T G G A T A C T A A A T G T T G T C T T A T T T T T T T C	Majority
	4510 4520 4530 4540 4550	
4501	C A A T T A T A G T G A C A A G C A A A A T G G A T A C T A A A T G T T G T C T T A T T T T T T T C	coh1_al2.seq
4501	C A A T T A T A G T G A C A A G C A A A A T G G A T A C T A A A T G T T G T C T T A T T T T T T T C	a909_al2.seq

FIGURE 21F



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      ATATATTTTAAATCTGTACCACTTTGCTAGCCCATCTTATTAAGAACGTA Majority
      4560      4570      4580      4590      4600
4551 ATATATTTTAAATCTGTACCACTTTGCTAGCCCATCTTATTAAGAACGTA coh1_al2.seq
4551 ATATATTTTAAATCTGTACCACTTTGCTAGCCCATCTTATTAAGAACGTA a909_al2.seq

      AACGACGACGAGCAACAAGCAGGATACCTGCTCCTATTACTAAAATTGCA Majority
      4610      4620      4630      4640      4650
4601 AACGACGACGAGCAACAAGCAGGATACCTGCTCCTATTACTAAAATTGCA coh1_al2.seq
4601 AACGACGACGAGCAACAAGCAGGATACCTGCTCCTATTACTAAAATTGCA a909_al2.seq

      CCTATAATGTAGAAAATTGTTGTACCAATACCACCTGTTGAAGGCAACTC Majority
      4660      4670      4680      4690      4700
4651 CCTATAATGTAGAAAATTGTTGTACCAATACCACCTGTTGAAGGCAACTC coh1_al2.seq
4651 CCTATAATGTAGAAAATTGTTGTACCAATACCACCTGTTGAAGGCAACTC a909_al2.seq

      AGTACCTTTGTTATTTTCAACAGTTGGGTTAACTAAAAGGTTATCTGAAT Majority
      4710      4720      4730      4740      4750
4701 AGTACCTTTGTTATTTTCAACAGTTGGGTTAACTAAAAGGTTATCTGAAT coh1_al2.seq
4701 AGTACCTTTGTTATTTTCAACAGTTGGGTTAACTAAAAGGTTATCTGAAT a909_al2.seq

      TAGTCGTATCAGTGGCTCCATCTCCTAAAATAACCTTCTGAGAGTTATCT Majority
      4760      4770      4780      4790      4800
4751 TAGTCGTATCAGTGGCTCCATCTCCTAAAATAACCTTCTGAGAGTTATCT coh1_al2.seq
4751 TAGTCGTATCAGTGGCTCCATCTCCTAAAATAACCTTCTGAGAGTTATCT a909_al2.seq

      AACAAATTGTAACCTAAGGGAGCCTTTTTCTCAACTAGATAGTATGTACC Majority
      4810      4820      4830      4840      4850
4801 AACAAATTGTAACCTAAGGGAGCCTTTTTCTCAACTAGATAGTATGTACC coh1_al2.seq
4801 AACAAATTGTAACCTAAGGGAGCCTTTTTCTCAACTAGATAGTATGTACC a909_al2.seq

      TTCTTTCAAGCCTGTAATGGTAATTATACCATCTGCTCCTGTTGTATATT Majority
      4860      4870      4880      4890      4900
4851 TTCTTTCAAGCCTGTAATGGTAATTATACCATCTGCTCCTGTTGTATATT coh1_al2.seq
4851 TTCTTTCAAGCCTGTAATGGTAATTATACCATCTGCTCCTGTTGTATATT a909_al2.seq

      CTGTTGCATTAGCTTCTGTGCCCCATTCAACGTTATTTGTATCGTTAAAG Majority
      4910      4920      4930      4940      4950
4901 CTGTTGCATTAGCTTCTGTGCCCCATTCAACGTTATTTGTATCGTTAAAG coh1_al2.seq
4901 CTGTTGCATTAGCTTCTGTGCCCCATTCAACGTTATTTGTATCGTTAAAG a909_al2.seq

      TTTAGAAATTGACCCGTAGCATTCTTTAAACAAATATAGCACCTTGTAAT Majority
      4960      4970      4980      4990      5000
4951 TTTAGAAATTGACCCGTAGCATTCTTTAAACAAATATAGCACCTTGTAAT coh1_al2.seq
4951 TTTAGAAATTGACCCGTAGCATTCTTTAAACAAATATAGCACCTTGTAAT a909_al2.seq

      TGAAGCTTTTGTGGAACCATCAATTTTTTTTATAGTAATTTGACCATCCC Majority
      5010      5020      5030      5040      5050
5001 TGAAGCTTTTGTGGAACCATCAATTTTTTTTATAGTAATTTGACCATCCC coh1_al2.seq
5001 TGAAGCTTTTGTGGAACCATCAATTTTTTTTATAGTAATTTGACCATCCC a909_al2.seq

      TCACCTGTTACTTTTTGACCTGGGTCATCATTGCTAGTATTGGGGTTGATG Majority
      5060      5070      5080      5090      5100
5051 TCACCTGTTACTTTTTGACCTGGGTCATCATTGCTAGTATTGGGGTTGATG coh1_al2.seq
5051 TCACCTGTTACTTTTTGACCTGGGTCATCATTGCTAGTATTGGGGTTGATG a909_al2.seq

      GTCGCAATGTTTGTATTTTCTGGTAAATCAGCTGAACCTGGTTLTACCTCC Majority
      5110      5120      5130      5140      5150
5101 GTCGCAATGTTTGTATTTTCTGGTAAATCAGCTGAACCTGGTTLTACCTCC coh1_al2.seq
5101 GTCGCAATGTTTGTATTTTCTGGTAAATCAGCTGAACCTGGTTLTACCTCC a909_al2.seq

      ACTCTTTAATACTCCTGTATAAGTGACTGTGATTGTATTTATTCCCTTAT Majority
      5160      5170      5180      5190      5200
5151 ACTCTTTAATACTCCTGTATAAGTGACTGTGATTGTATTTATTCCCTTAT coh1_al2.seq
5151 ACTCTTTAATACTCCTGTATAAGTGACTGTGATTGTATTTATTCCCTTAT a909_al2.seq

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FIGURE 21G

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      A A A A A A A G T C A T C A T T A G C T C C A A T T T G A G T A T T T C C G G T T G G A G T A T T G Majority
      5210      5220      5230      5240      5250
5201 A A A A A A A G T C A T C A T T A G C T C C A T T T T G A G T A T T T C C G G T T G G A G T A T T G coh1_al2.seq
5201 A A A A A A A G T C A T C A T T A G C T C C A T T T T G A G T A T T T C C G G T T G G A G T A T T G a909_al2.seq

      G T A G C T G C C C A C G G A A T A G T A A T C G T G A A A T T A T T A T T T T C C T C T A A C A G Majority
      5260      5270      5280      5290      5300
5251 G T A G C T G C C C A C G G A A T A G T A A T C G T G A A A T T A T T A T T T T C C T C T A A C A G coh1_al2.seq
5251 G T A G C T G C C C A C G G A A T A G T A A T C G T G A A A T T A T T A T T T T C C T C T A A C A G a909_al2.seq

      G T T A T A C T T C C C A G T T G C T T T T T C C G A A C C T T G A G T T A G A G T T G T A A T A T Majority
      5310      5320      5330      5340      5350
5301 G T T A T A C T T C C C A G T T G C T T T T T C C G A A C C T T G A G T T A G A G T T G T A A T A T coh1_al2.seq
5301 G T T A T A C T T C C C A G T T G C T T T T T C C G A A C C T T G A G T T A G A G T T G T A A T A T a909_al2.seq

      T C C C T G A T C C A T C A G T A A T A G T T A C T T C A T A A G A T C C T T C G T T C A A A T C A Majority
      5360      5370      5380      5390      5400
5351 T C C C T G A T C C A T C A G T A A T A G T T A C T T C A T A A G A T C C T T C G T T C A A A T C A coh1_al2.seq
5351 T C C C T G A T C C A T C A G T A A T A G T T A C T T C A T A A G A T C C T T C G T T C A A A T C A a909_al2.seq

      A C T A C A G A A G C A G A T G G C A T A G T A T C C T T T A T A A C A T A T T G A T A C A C T T T Majority
      5410      5420      5430      5440      5450
5401 A C T A C A G A A G C A G A T G G C A T A G T A T C C T T T A T A A C A T A T T G A T A C A C T T T coh1_al2.seq
5401 A C T A C A G A A G C A G A T G G C A T A G T A T C C T T T A T A A C A T A T T G A T A C A C T T T a909_al2.seq

      T T C T G T A C C A T G A T A A T T G A C T G C A T T C T T A T A A G T A A T A G T A T A T T T G A Majority
      5460      5470      5480      5490      5500
5451 T T C T G T A C C A T G A T A A T T G A C T G C A T T C T T A T A A G T A A T A G T A T A T T T G A coh1_al2.seq
5451 T T C T G T A C C A T G A T A A T T G A C T G C A T T C T T A T A A G T A A T A G T A T A T T T G A a909_al2.seq

      C T G T A T C A C C A A C C G A G T A C G T T T T T T G A T C T A C A G T T T T T C C A C C A C C A Majority
      5510      5520      5530      5540      5550
5501 C T G T A T C A C C A A C C G A G T A C G T T T T T T G A T C T A C A G T T T T T C C A C C A C C A coh1_al2.seq
5501 C T G T A T C A C C A A C C G A G T A C G T T T T T T G A T C T A C A G T T T T T C C A C C A C C A a909_al2.seq

      T C T C C C C A T G T C G C A T C A G T A T T C T T T T C A T G A A T A G T A G C A T T T G G A G T Majority
      5560      5570      5580      5590      5600
5551 T C T C C C C A T G T C G C A T C A G T A T T C T T T T C A T G A A T A G T A G C A T T T G G A G T coh1_al2.seq
5551 T C T C C C C A T G T C G C A T C A G T A T T C T T T T C A T G A A T A G T A G C A T T T G G A G T a909_al2.seq

      T A C A G A T G T A A C C A T A A T T A C A G C T C C A T T A T T A A C A G T G C T A G A A A C A T Majority
      5610      5620      5630      5640      5650
5601 T A C A G A T G T A A C C A T A A T T A C A G C T C C A T T A T T A A C A G T G C T A G A A A C A T coh1_al2.seq
5601 T A C A G A T G T A A C C A T A A T T A C A G C T C C A T T A T T A A C A G T G C T A G A A A C A T a909_al2.seq

      A A T A A T A T C C A T A T T G G G A A A C A T T A A T A A C C T C A G T A C C A T C A T T A T T T Majority
      5660      5670      5680      5690      5700
5651 A A T A A T A T C C A T A T T G G G A A A C A T T A A T A A C C T C A G T A C C A T C A T T A T T T coh1_al2.seq
5651 A A T A A T A T C C A T A T T G G G A A A C A T T A A T A A C C T C A G T A C C A T C A T T A T T T a909_al2.seq

      G A C T C A G T A A C A G T G G A A A C T G G T G T A G T A T T A G C T G A T A T A G A T T T A G C Majority
      5710      5720      5730      5740      5750
5701 G A C T C A G T A A C A G T G G A A A C T G G T G T A G T A T T A G C T G A T A T A G A T T T A G C coh1_al2.seq
5701 G A C T C A G T A A C A G T G G A A A C T G G T G T A G T A T T A G C T G A T A T A G A T T T A G C a909_al2.seq

      C C A T G T C G C A A T C T C A T T T G C T G A C G C A G T A T C T T T T T A G T T A C A T A T G Majority
      5760      5770      5780      5790      5800
5751 C C A T G T C G C A A T C T C A T T T G C T G A C G C A G T A T C T T T T T A G T T A C A T A T G coh1_al2.seq
5751 C C A T G T C G C A A T C T C A T T T G C T G A C G C A G T A T C T T T T T A G T T A C A T A T G a909_al2.seq

      T T C T C C C T C C A T T A G T A G T T G T C G T A A A A A G A G A A T T A A A A T C A G T T G A A Majority
      5810      5820      5830      5840      5850
5801 T T C T C C C T C C A T T A G T A G T T G T C G T A A A A A G A G A A T T A A A A T C A G T T G A A coh1_al2.seq
5801 T T C T C C C T C C A T T A G T A G T T G T C G T A A A A A G A G A A T T A A A A T C A G T T G A A a909_al2.seq

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FIGURE 21H



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G C T T A A A C T C A G C T T C T T T A C C T T T G A G G A A T T A A A T A A G A A G C T C C A T C Majority
5860      5870      5880      5890      5900
5851 G C T T T A T A C T C A G C T T C T T T A C C T T G A G G A A T T A A A T A A G A A G C T C C A T C coh1_al2.seq
5851 G C T T T A T A C T C A G C T T C T T T A C C T T G A G G A A T T A A A T A A G A A G C T C C A T C a909_al2.seq

T T T A T T C G A A T C A G A T A C A T T T G C A T T A T C T A T T T T C T G C A T C A A A A A C T T Majority
5910      5920      5930      5940      5950
5901 T T T A T T C G A A T C A G A T A C A T T T G C A T T A T C T A T T T T C T G C A T C A A A A A C T T coh1_al2.seq
5901 T T T A T T C G A A T C A G A T A C A T T T G C A T T A T C T A T T T T C T G C A T C A A A A A C T T a909_al2.seq

T G T A T G C T T T A T A G G T T G C G C C T T T T T G A G T A T C T T G A A C T G T A A T T G T C Majority
5960      5970      5980      5990      6000
5951 T A T A T G C T T T A T A G G T T G C G C C T T T T T G A G T A T C T T G A A C T G T A A T T G T C coh1_al2.seq
5951 T G T A T G C T T T A T A G G T T G C G C C T T T T T G A G T A T C T T G A A C T G T A A T T G T C a909_al2.seq

C C T G T C T C A G C G G C A A A A G C T A T C G G C G T A A C T G G T G A T A C A G C C A T A C C Majority
6010      6020      6030      6040      6050
6001 C C T G T C T C A G C G G C A A A A G C T A T C G G C G T A A C T G G T G A T A C A G C C A T A C C coh1_al2.seq
6001 C C T G T C T C A G C G G C A A A A G C T A T C G G C G T A A C T G G T G A T A C A G C C A T A C C a909_al2.seq

A A A T G C T A A A C T C G C C A C T A A C A G C G A T T G A A T C A T T T T C T T T T T C A T T G Majority
6060      6070      6080      6090      6100
6051 A A A T G C T A A A C T C G C C A C T A A C A G C G A T T G A A T C A T T T T C T T T T T C A T T G coh1_al2.seq
6051 A A A T G C T A A A C T C G C C A C T A A C A G C G A T T G A A T C A T T T T C T T T T T C A T T G a909_al2.seq

A A A T C T T T C T C C T A A A A T C A T A T T G A T G A A T G A T T A A T T C A T A T T T T T T Majority
6110      6120      6130      6140      6150
6101 A A A T C T T T C T C C T A A A A T C A T A T T G A T G A A T G A T T A A T T C A T A T T T T T T coh1_al2.seq
6101 A A A T C T T T C T C C T A A A A T C A T A T T G A T G A A T G A T T A A T T C A T A T T T T T T a909_al2.seq

T C G A T A G T A T A A T A T T A A T C C T G A T G G T A G A G C T A A A G C T A A A C C A A C T A Majority
6160      6170      6180      6190      6200
6151 T C G A T A G T A T A A T A T T A A T C C T G A T G G T A G A G C T A A A G C T A A A C C A A C T A coh1_al2.seq
6151 T C G A T A G T A T A A T A T T A A T C C T G A T G G T A G A G C T A A A G C T A A A C C A A C T A a909_al2.seq

G G A T A T A A A T G T G T G T T C C A A T A C C T C C A G T A C T A G G C A A T T C T G T T C C T Majority
6210      6220      6230      6240      6250
6201 G G A T A T A A A T G T G T G T T C C A A T A C C T C C A G T A C T A G G C A A T T C T G T T C C T coh1_al2.seq
6201 G G A T A T A A A T G T G T G T T C C A A T A C C T C C A G T A C T A G G C A A T T C T G T T C C T a909_al2.seq

T T A C T G T T A G T A A T T T T A A A A G T A T A T A C T G T A C T T C C A T C T A C T A A A T T Majority
6260      6270      6280      6290      6300
6251 T T A C T G T T A G T A A T T T T A A A A G T A T A T A C T G T A C T T C C A T C T A C T A A A T T coh1_al2.seq
6251 T T A C T G T T A G T A A T T T T A A A A G T A T A T A C T G T A C T T C C A T C T A C T A A A T T a909_al2.seq

C T C T T T T A T T G G T G T C G C A T T A T T A C C A T T T T G T T C A A A G G T A A C T C C C G Majority
6310      6320      6330      6340      6350
6301 C T C T T T T A T T G G T G T C G C A T T A T T A C C A T T T T G T T C A A A G G T A A C T C C C G coh1_al2.seq
6301 C T C T T T T A T T G G T G T C G C A T T A T T A C C A T T T T G T T C A A A G G T A A C T C C C G a909_al2.seq

T A G A A A T C A C T A A T A C T G A T A T A T C A T T T T T A G G T A G T A G G T A C C C T G G A Majority
6360      6370      6380      6390      6400
6351 T A G A A A T C A C T A A T A C T G A T A T A T C A T T T T T A G G T A G T A G G T A C C C T G G A coh1_al2.seq
6351 T A G A A A T C A C T A A T A C T G A T A T A T C A T T T T T A G G T A G T A G G T A C C C T G G A a909_al2.seq

G G G G C C T T T G T C T C T G T T A G G T A G T A T T T T C C T A C T G G C A A A C T G A G G T A Majority
6410      6420      6430      6440      6450
6401 G G G G C C T T T G T C T C T G T T A G G T A G T A T T T T C C T A C T G G C A A A C T G A G G T A coh1_al2.seq
6401 G G G G C C T T T G T C T C T G T T A G G T A G T A T T T T C C T A C T G G C A A A C T G A G G T A a909_al2.seq

G T T A T T A G C A T C C A C T A A T A A C A A G C C T T T A T C G T T T G T C A C C A G C C C T G Majority
6460      6470      6480      6490      6500
6451 G T T A T T A G C A T C C A C T A A T A A C A A G C C T T T A T C G T T T G T C A C C A G C C C T G coh1_al2.seq
6451 G T T A T T A G C A T C C A C T A A T A A C A A G C C T T T A T C G T T T G T C A C C A G C C C T G a909_al2.seq

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FIGURE 211

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AATAAATAAGGATGTGAAGCTTTATTCCATTAGCATCTGATTTCATAAATA Majority
6510 6520 6530 6540 6550
6501 AATACATAGGATGTGAAGCTTTATTCCATTAGCATCTGATTTCATAAATA coh1_al2.seq
6501 AATACATAGGATGTGAAGCTTTATTCCATTAGCATCTGATTTCATAAATA a909_al2.seq

TCAAAAACTGCACCTGCTAAAAAATTATTATCATTTTTCGACATTAACTTT Majority
6560 6570 6580 6590 6600
6551 TCAAAAACTGCACCTGCTAAAAAATTATTATCATTTTTCGACATTAACTTT coh1_al2.seq
6551 TCAAAAACTGCACCTGCTAAAAAATTATTATCATTTTTCGACATTAACTTT a909_al2.seq

CTGTAGTCGTACTTTTTCCTTGATACGTGTATTGGTAAAGCTAATATCTA Majority
6610 6620 6630 6640 6650
6601 CTGTAGTCGTACTTTTTCCTTGATACGTGTATTGGTAAAGCTAATATCTA coh1_al2.seq
6601 CTGTAGTCGTACTTTTTCCTTGATACGTGTATTGGTAAAGCTAATATCTA a909_al2.seq

CGTCTCCTGAAACTGTCAGGGATTGTAAGCCGGTAGCATCATAAGTTTTA Majority
6660 6670 6680 6690 6700
6651 CGTCTCCTGAAACTGTCAGGGATTGTAAGCCGGTAGCATCATAAGTTTTA coh1_al2.seq
6651 CGTCTCCTGAAACTGTCAGGGATTGTAAGCCGGTAGCATCATAAGTTTTA a909_al2.seq

TCAGCTTCACCAGTTGCTAGATTTTTTTCTGTAAATTGACTCAGATACTTT Majority
6710 6720 6730 6740 6750
6701 TCAGCTTCACCAGTTGCTAGATTTTTTTCTGTAAATTGACTCAGATACTTT coh1_al2.seq
6701 TCAGCTTCACCAGTTGCTAGATTTTTTTCTGTAAATTGACTCAGATACTTT a909_al2.seq

AAATTCATCGTAGGCTTGTTCATCTATTGATATAGAAGTTCCATAAAGGTA Majority
6760 6770 6780 6790 6800
6751 AAATTCATCGTAGGCTTGTTCATCTATTGATATAGAAGTTCCATAAAGGTA coh1_al2.seq
6751 AAATTCATCGTAGGCTTGTTCATCTATTGATATAGAAGTTCCATAAAGGTA a909_al2.seq

CTTTAAATTGCTTAGTCTGACCATCTCTCAGCGGAAAATTCTCTTGTTCG Majority
6810 6820 6830 6840 6850
6801 CTTTAAATTGCTTAGTCTGACCATCTCTCAGCGGAAAATTCTCTTGTTCG coh1_al2.seq
6801 CTTTAAATTGCTTAGTCTGACCATCTCTCAGCGGAAAATTCTCTTGTTCG a909_al2.seq

AACGTTTCACTTGGATTAAACAAGAAGTCTTTTCGTCTTATCTTCATCTAG Majority
6860 6870 6880 6890 6900
6851 AACGTTTCACTTGGATTAAACAAGAAGTCTTTTCGTCTTATCTTCATCTAG coh1_al2.seq
6851 AACGTTTCACTTGGATTAAACAAGAAGTCTTTTCGTCTTATCTTCATCTAG a909_al2.seq

TCCAACGACAGTTTTTACTTACTCTGACGGTGTATTCTTTAGGTTGCCAAA Majority
6910 6920 6930 6940 6950
6901 TCCAACGACAGTTTTTACTTACTCTGACGGTGTATTCTTTAGGTTGCCAAA coh1_al2.seq
6901 TCCAACGACAGTTTTTACTTACTCTGACGGTGTATTCTTTAGGTTGCCAAA a909_al2.seq

CAGCATATAAGGTATTTTGTTCGATCAGGGTTGTTATCAATACCTATTGAT Majority
6960 6970 6980 6990 7000
6951 CAGCATATAAGGTATTTTGTTCGATCAGGGTTGTTATCAATACCTATTGAT coh1_al2.seq
6951 CAGCATATAAGGTATTTTGTTCGATCAGGGTTGTTATCAATACCTATTGAT a909_al2.seq

TGACCTGCTGTAAATTCCACACGTCCTGTATCAGCTAAATCCTTATCATG Majority
7010 7020 7030 7040 7050
7001 TGACCTGCTGTAAATTCCACACGTCCTGTATCAGCTAAATCCTTATCATG coh1_al2.seq
7001 TGACCTGCTGTAAATTCCACACGTCCTGTATCAGCTAAATCCTTATCATG a909_al2.seq

ATGCCAACCATAAAGGTTGTAACCTGTCCTTGTAAAGTATTGGTTTTTCAG Majority
7060 7070 7080 7090 7100
7051 ATGCCAACCATAAAGGTTGTAACCTGTCCTTGTAAAGTATTGGTTTTTCAG coh1_al2.seq
7051 ATGCCAACCATAAAGGTTGTAACCTGTCCTTGTAAAGTATTGGTTTTTCAG a909_al2.seq

GAATTGTAGTTGTGCTATTCAACTCCATACCGCGGTGTCTCTACTTGTGTT Majority
7110 7120 7130 7140 7150
7101 GAATTGTAGTTGTGCTATTCAACTCCATACCGCGGTGTCTCTACTTGTGTT coh1_al2.seq
7101 GAATTGTAGTTGTGCTATTCAACTCCATACCGCGGTGTCTCTACTTGTGTT a909_al2.seq

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FIGURE 21J



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ACCACATTACCCTTTCTACTCTAGTACCACCGTTACCATTGTATTTGAT Majority  
7160 7170 7180 7190 7200  
7151 ACCACATTACCCTTTCTACTCTAGTACCACCGTTACCATTGTATTTGAT coh1\_al2.seq  
7151 ACCACATTACCCTTTCTACTCTAGTACCACCGTTACCATTGTATTTGAT a909\_al2.seq  
TCAGGTATCTTCTAATTTGATATCTCCTACTGGAATAATGACAGGTTTTA Majority  
7210 7220 7230 7240 7250  
7201 TCAGGTATCTTCTAATTTGATATCTCCTACTGGAATAATGACAGGTTTTA coh1\_al2.seq  
7201 TCAGGTATCTTCTAATTTGATATCTCCTACTGGAATAATGACAGGTTTTA a909\_al2.seq  
TGGTGATATTTTTTATTAGCATCTGCTAAATGGGCGTCAATATCAATGGAA Majority  
7260 7270 7280 7290 7300  
7251 TGGTGATATTTTTTATTAGCATCTGCTAAATGGGCGTCAATATCAATGGAA coh1\_al2.seq  
7251 TGGTGATATTTTTTATTAGCATCTGCTAAATGGGCGTCAATATCAATGGAA a909\_al2.seq  
TCATATGGGTTATAAATTTTACCATTGTACCACCAGCCACGGAAACGATA Majority  
7310 7320 7330 7340 7350  
7301 TCATATGGGTTATAAATTTTACCATTGTACCACCAGCCACGGAAACGATA coh1\_al2.seq  
7301 TCATATGGGTTATAAATTTTACCATTGTACCACCAGCCACGGAAACGATA a909\_al2.seq  
GCCATCTGGCATTGTCCGACGCTCTCAGTAAGGCTGAATGGGAGCCATCGT Majority  
7360 7370 7380 7390 7400  
7351 GCCATCTGGCATTGTCCGACGCTCTCAGTAAGGCTGAATGGGAGCCATCGT coh1\_al2.seq  
7351 GCCATCTGGCATTGTCCGACGCTCTCAGTAAGGCTGAATGGGAGCCATCGT a909\_al2.seq  
CATAGGAATCAGGGTCAAGTACGCTCATTACTTGTTTTGTAAATTGCTGACCA Majority  
7410 7420 7430 7440 7450  
7401 CATAGGAATCAGGGTCAAGTACGCTCATTACTTGTTTTGTAAATTGCTGACCA coh1\_al2.seq  
7401 CATAGGAATCAGGGTCAAGTACGCTCATTACTTGTTTTGTAAATTGCTGACCA a909\_al2.seq  
GAAGCATCCAATGCTGGCTTTCCATCTGTACCAACAGCATCATTGCTGTA Majority  
7460 7470 7480 7490 7500  
7451 GAAGCATCCAATGCTGGCTTTCCATCTGTACCAACAGCATCATTGCTGTA coh1\_al2.seq  
7451 GAAGCATCCAATGCTGGCTTTCCATCTGTACCAACAGCATCATTGCTGTA a909\_al2.seq  
TATAATATGATAATCTCCAGCCTTTTCGCCAAATAGCTCTTAAATTGATAT Majority  
7510 7520 7530 7540 7550  
7501 TATAATATGATAATCTCCAGCCTTTTCGCCAAATAGCTCTTAAATTGATAT coh1\_al2.seq  
7501 TATAATATGATAATCTCCAGCCTTTTCGCCAAATAGCTCTTAAATTGATAT a909\_al2.seq  
CTTGAGTTACAGCACCTGAAAAGTTATAAGGTCTAATACTACCATCTGGA Majority  
7560 7570 7580 7590 7600  
7551 CTTGAGTTACAGCACCTGAAAAGTTATAAGGTCTAATACTACCATCTGGA coh1\_al2.seq  
7551 CTTGAGTTACAGCACCTGAAAAGTTATAAGGTCTAATACTACCATCTGGA a909\_al2.seq  
TTAACATAAATACCAACCGACTAATTTGTAAAGCGTCTTTTACGTACTTGTAA Majority  
7610 7620 7630 7640 7650  
7601 TTAACATAAATACCAACCGACTAATTTGTAAAGCGTCTTTTACGTACTTGTAA coh1\_al2.seq  
7601 TTAACATAAATACCAACCGACTAATTTGTAAAGCGTCTTTTACGTACTTGTAA a909\_al2.seq  
CTTAGTTGTGTATCAACATTTGAGAGACTAGTATCTGTCTGTATAAATAGG Majority  
7660 7670 7680 7690 7700  
7651 CTTAGTTGTGTATCAACATTTGAGAGACTAGTATCTGTCTGTATAAATAGG coh1\_al2.seq  
7651 CTTAGTTGTGTATCAACATTTGAGAGACTAGTATCTGTCTGTATAAATAGG a909\_al2.seq  
CATCTTTAGTTGAGTCCGGATCTTTATCTCGTGAATCATACTTATAATAA Majority  
7710 7720 7730 7740 7750  
7701 CATCTTTAGTTGAGTCCGGATCTTTATCTCGTGAATCATACTTATAATAA coh1\_al2.seq  
7701 CATCTTTAGTTGAGTCCGGATCTTTATCTCGTGAATCATACTTATAATAA a909\_al2.seq  
TATGTACCTGAAGGATCTTTGGATATAATCCCTTGTAATATCTGTATAATC Majority  
7760 7770 7780 7790 7800  
7751 TATGTACCTGAAGGATCTTTGGATATAATCCCTTGTAATATCTGTATAATC coh1\_al2.seq  
7751 TATGTACCTGAAGGATCTTTGGATATAATCCCTTGTAATATCTGTATAATC a909\_al2.seq

FIGURE 21K

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CGGAAATACGATCACCATAATGCAAATCTAAATAGGTATCATCTGTTTTTG Majority  
7810 7820 7830 7840 7850  
7801 CGGAAATACGATCACCATAATGCAAATCTAAATAGGTATCATCTGTTTTTG coh1\_al2.seq  
7801 CGGAAATACGATCACCATAATGCAAAGTCTAAATAGGTATCATCTGTTTTTG a909\_al2.seq  
ATAATTGGCCTCCGTTTGGATCAATATTGACACGATATGTTACCTTTTGC Majority  
7860 7870 7880 7890 7900  
7851 ATAATTGGCCTCCGTTTGGATCAATATTGACACGATATGTTACCTTTTGC coh1\_al2.seq  
7851 ATAATTGGCCTCCGTTTGGATCAATATTGACACGATATGTTACCTTTTGC a909\_al2.seq  
CAACCTGCATAGACTTTTAACATCATGAGGAGGCATAGTCGTGTTAAAGTC Majority  
7910 7920 7930 7940 7950  
7901 CAACCTGCATAGACTTTTAACATCATGAGGAGGCATAGTCGTGTTAAAGTC coh1\_al2.seq  
7901 CAACCTGCATAGACTTTTAACATCATGAGGAGGCATAGTCGTGTTAAAGTC a909\_al2.seq  
AAATACTTGTGTTTGTGCTTGGTCTTTATACCATTTACCATCCCAAACAT Majority  
7960 7970 7980 7990 8000  
7951 AAATACTTGTGTTTGTGCTTGGTCTTTATACCATTTACCATCCCAAACAT coh1\_al2.seq  
7951 AAATACTTGTGTTTGTGCTTGGTCTTTATACCATTTACCATCCCAAACAT a909\_al2.seq  
ACCCTGGTCGACTAGGTTTAGGTTGAACCGTTGTCGTATCGGGGGGCATAA Majority  
8010 8020 8030 8040 8050  
8001 ACCCTGGTCGACTAGGTTTAGGTTGAACCGTTGTCGTATCGGGGGGCATAA coh1\_al2.seq  
8001 ACCCTGGTCGACTAGGTTTAGGTTGAACCGTTGTCGTATCGGGGGGCATAA a909\_al2.seq  
GAGGACAAATTTTGTCTCATATAGAACATCCTTTACTGGAAAATTAGGAAG Majority  
8060 8070 8080 8090 8100  
8051 GAGGACAAATTTTGTCTCATATAGAACATCCTTTACTGGAAAATTAGGAAG coh1\_al2.seq  
8051 GAGGACAAATTTTGTCTCATATAGAACATCCTTTACTGGAAAATTAGGAAG a909\_al2.seq  
CTCTGTATTATCAAGCGGATCTAAATATTTAATCTTGTATGAATTACGTT Majority  
8110 8120 8130 8140 8150  
8101 CTCTGTATTATCAAGCGGATCTAAATATTTAATCTTGTATGAATTACGTT coh1\_al2.seq  
8101 CTCTGTATTATCAAGCGGATCTAAATATTTAATCTTGTATGAATTACGTT a909\_al2.seq  
CATACCATAACCACTAAGTTCAAATAATCTTTGTGCTAGTCTCCATATTTA Majority  
8160 8170 8180 8190 8200  
8151 CATACCATAACCACTAAGTTCAAATAATCTTTGTGCTAGTCTCCATATTTA coh1\_al2.seq  
8151 CATACCATAACCACTAAGTTCAAATAATCTTTGTGCTAGTCTCCATATTTA a909\_al2.seq  
TCGTAGTATTTCATCTGCGATTGGCACTTTTGTGTTTTTGCACCTCGTTTGTCT Majority  
8210 8220 8230 8240 8250  
8201 TCGTAGTATTTCATCTGCGATTGGCACTTTTGTGTTTTTGCACCTCGTTTGTCT coh1\_al2.seq  
8201 TCGTAGTATTTCATCTGCGATTGGCACTTTTGTGTTTTTGCACCTCGTTTGTCT a909\_al2.seq  
TGGGTTCTGATCAAAATAGGTAATTATCTGGATATAAGCTTTGATAGTATT Majority  
8260 8270 8280 8290 8300  
8251 TGGGTTCTGATCAAAATAGGTAATTATCTGGATATAAGCTTTGATAGTATT coh1\_al2.seq  
8251 TGGGTTCTGATCAAAATAGGTAATTATCTGGATATAAGCTTTGATAGTATT a909\_al2.seq  
TAACATTAAATCCTAGGCTATTTTCTGTAAGGTAATTCCTCTGCTCCA Majority  
8310 8320 8330 8340 8350  
8301 TAACATTAAATCCTAGGCTATTTTCTGTAAGGTAATTCCTCTGCTCCA coh1\_al2.seq  
8301 TAACATTAAATCCTAGGCTATTTTCTGTAAGGTAATTCCTCTGCTCCA a909\_al2.seq  
GCACCTCCCTGCTGCTGCTAAAGAGTATTGCCCATCTAGTCCTTGTTT Majority  
8360 8370 8380 8390 8400  
8351 GCACCTCCCTGCTGCTGCTAAAGAGTATTGCCCATCTAGTCCTTGTTT coh1\_al2.seq  
8351 GCACCTCCCTGCTGCTGCTAAAGAGTATTGCCCATCTAGTCCTTGTTT a909\_al2.seq  
GTAGAACGGATAAATTTGAATTCTCTTCCCTTTTGGATAGAGTTTATTT Majority  
8410 8420 8430 8440 8450  
8401 GTAGAACGGATAAATTTGAATTCTCTTCCCTTTTGGATAGAGTTTATTT coh1\_al2.seq  
8401 GTAGAACGGATAAATTTGAATTCTCTTCCCTTTTGGATAGAGTTTATTT a909\_al2.seq

FIGURE 21L



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CATCTGCAATTTGCTGTAATGTAAGTCCATGGGTAGTATGAACCTCACCCAAA Majority
      8460      8470      8480      8490      8500
8451 CATCTGCAATCAACGCTAT...CATTAGGTAAAATGAACCTCACCCAAA coh1_al2.seq
8451 CATAGGATTTCCTGCTGTAAGTACTCCAATGGGAAGTATAAAGTCTCACCCAAA a909_al2.seq

TAACTCATTCCTTATGTTCCAGTTTGGTTATTTCCAACATTGGTTAGGTA Majority
      8510      8520      8530      8540      8550
8495 TAACTCATTCCTATAAGGTTCCAAGTTGGTTATTTCCAACATTGGCTAATA coh1_al2.seq
8501 TAACTCATTCCTGATAGTCCAGTTGAA...GTTAGGTA a909_al2.seq

ACGCCATGCACCTGTCTTCCATTGATAGCCATTGCGCGGCTAAGGTTGTAC Majority
      8560      8570      8580      8590      8600
8545 ACGCCATGCACCGCTCTTCCATTGATAACCATTAAGCGGCCTAAGGTTGTAC coh1_al2.seq
8536 ACGCCAAGCACCTGTCTTCCAATGATAGCCATTGCGAGCTAAAGTGTGTAC a909_al2.seq

CGTATAGTCTCTGTGTAGGTTTCCGGCATCTGATGCTCTAGTTCTAGGAATC Majority
      8610      8620      8630      8640      8650
8595 CGTAAGATCTCTGTGTAGGTAAGTATCAGCATCACAAGCTCTCAATTATAGGAATA coh1_al2.seq
8586 CATATAGTCTCAGTATAGGTTTCCGGCATATGATGCACTAGTCTCAAGGAATAC a909_al2.seq

GTAGTATTTTGGTAATGAATCTCCGAGTAGCCCTTTTTTGCATAATTTTAT Majority
      8660      8670      8680      8690      8700
8644 GTAAATATTTTGGCAAGGAATATCCG...CCATATTTTGCATAATTTTAT coh1_al2.seq
8636 GAAGAAGTTTGGATAATAAACTCTAGAGTAGCCCTTTTGCATAATCTTAT a909_al2.seq

TGTGATGAGTTTCTATCATAATAAACATTAACGACACTTGAACCATCGT Majority
      8710      8720      8730      8740      8750
8689 TGTGATGAGTTTCTATCATAATAAACATTAACGACACTTGAACCATCGT coh1_al2.seq
8686 AGTGAAGATTTCTATCATAATAAACATTAACGACACTTGAACCATCGT a909_al2.seq

CTTTTATCATGACAGAAGTTTCTGTCTCTCGTATTATTAACTTTAAAGCCA Majority
      8760      8770      8780      8790      8800
8739 CTTTTATCATGACAGAAGTTTCTGTCTCTCGTATTATTAACTTTAAAGCCA coh1_al2.seq
8736 CTTTTATCATGACAGAAGTTTCTGTCTCTCGTATTATTAACTTTAAAGCCA a909_al2.seq

GTCCGTAGTTTTTTCATTAATATCTTGTGTGTAGCGTCTGATTAGATAA Majority
      8810      8820      8830      8840      8850
8789 GTCCGTAGTTTTATCATTAATATCTTGTGTGTAGCGTCTGATTAGATAA coh1_al2.seq
8786 GTCCGGAATTTTTCATTAATATCTTGTGTGTAGCGTCTGATTAGATAA a909_al2.seq

AGATAGGCCCTGATCGTGTACTTGGCCCTGCGTACTCATATGCTCTTTTGGC Majority
      8860      8870      8880      8890      8900
8839 AGATAGGCCCTGATCGTGTACTTGGCCCTGCGTACTCATATGCTCTTTTGGC coh1_al2.seq
8836 AGATAGGCCCTGATCGTGTACTTGGCCCTGCGTACTCATATGCTCTTTTGGC a909_al2.seq

CATCAGTAGCATTTTTATTATCCGTTGCTGATTGTTGCCAGTAGTTTATC Majority
      8910      8920      8930      8940      8950
8889 CATCAGTAGCATTTTTATTATCCGTTGCTGATTGTTGCCAGTAGTTTATC coh1_al2.seq
8886 CATCAGTAGCATTTTTATTATCCGTTGCTGATTGTTGCCAGTAGTTTATC a909_al2.seq

GTGTAGGTTGTTTGTGCGCGCGACCAATGTGCATATAGCGTCGTATCCTT Majority
      8960      8970      8980      8990      9000
8939 GTGTAGGTTGTTTGTGCGCGCGACCAATGTGCATATAGCGTCGTATCCTT coh1_al2.seq
8936 GTGTAGGTTGTTTGTGCGCGCGACCAATGTGCATATAGCGTCGTATCCTT a909_al2.seq

GGTCAAGACTTGATTAAAAATCAAAGGCTGTCCCAACCACTAGCAGCTGTGT Majority
      9010      9020      9030      9040      9050
8989 GGTCAAGACTTGATTAAAAATCAAAGGCTGTCCCAACCACTAGCAGCTGTGT coh1_al2.seq
8986 GGTCAAGACTTGATTAAAAATCAAAGGCTGTCCCAACCACTAGCAGCTGTGT a909_al2.seq

ACCAACCTTGCAAAAAGTATAACCTGGCCTTGTGATCATTAGGCTTAATT Majority
      9060      9070      9080      9090      9100
9039 ACCAACCTTGCAAAAAGTATAACCTGGCCTTGTGATCATTAGGCTTAATT coh1_al2.seq
9036 ACCAACCTTGCAAAAAGTATAACCTGGCCTTGTGATCATTAGGCTTAATT a909_al2.seq

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FIGURE 21M

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	G T C G A A G G A G G T T G C T C T G T T A A C A C A G G A C G A G G T G C A A T A T A G G T A A C	Majority
	9110 9120 9130 9140 9150	
9089	G T C G A A G C A G G T T G G C T G T T A A C A C A C G A C G A G G T G C A A T A T A G G T A A C	cohl_al2.seq
9086	G T C G A A G C A G G T T G G T C T G T T A A C A C A C G A C G A G G T G C A A T A T A G G T A A C	a909_al2.seq
	T C C T G T T G A T A A G T C G C C T G T G T T G A A T T C A A C A C C G T A C G A T T C T T T A A	Majority
	9160 9170 9180 9190 9200	
9139	T C C T G T T G A T A A G T C G C C T G T G T T G A A T T C A A C A C C G T A C G A T T C T T T A A	cohl_al2.seq
9136	T C C T G T T G A T A A G T C G C C T G T G T T G A A T T C A A C A C C G T A C G A T T C T T T A A	a909_al2.seq
	A G A C A G G A T A A A G A C T T A T G T C T G A A G A T A C A G G T A A T G C T T G A A T T T C T	Majority
	9210 9220 9230 9240 9250	
9189	A G A C A G G A T A A A G A C T T A T G T C T G A A G A T A C A G G T A A T G C T T G A A T T T C T	cohl_al2.seq
9186	A G A C A G G A T A A A G A C T T A T G T C T G A A G A T A C A G G T A A T G C T T G A A T T T C T	a909_al2.seq
	G A T T C A G A A A G T G G A G C T C C A T T T T G A G T T T T A G A C C A A C C T A C A A A T A A	Majority
	9260 9270 9280 9290 9300	
9239	G A T T C A G A A A G T G G A G C T C C A T T T T G A G T T T T A G A C C A A C C T A C A A A T A A	cohl_al2.seq
9236	G A T T C A G A A A G T G G A G C T C C A T T T T G A G T T T T A G A C C A A C C T A C A A A T A A	a909_al2.seq
	T A A T G T T G A A G T G G G T G G T G A A A C T T T A A A T G A A C T T A G A T C C G C G G T T C	Majority
	9310 9320 9330 9340 9350	
9289	T A A T G T T G A A G T G G G T G G T G A A A C T T T A A A T G A A C T T A A A T C C G C G G T T C	cohl_al2.seq
9286	T A A T G T T G A A G T G G G T G G T G A A A C T T T A A A T G A A C T T A G A T C C G C G G T T C	a909_al2.seq
	C G T T T C C T G A A C T A T C C A A T G G T A C T T G A T A A C G C T C C A A A A T A C T T T T A	Majority
	9360 9370 9380 9390 9400	
9339	C G T T T C C T G A A C T A T C C A A T G G T A C T T G A T A A C G C T C C A A A A T A C T T T T A	cohl_al2.seq
9336	C G T T T C C T G A A C T A T C C A A T G G T A C T T G A T A A C G C T C C A A A A T A C T T T T A	a909_al2.seq
	C C A G C A G A A T C A T C A T A T A A A G T T A C T G T C G C C A C T T T A C T C A T A A A T G G	Majority
	9410 9420 9430 9440 9450	
9389	C C A G C A G A A T C A T C A T A T A A A G T T A C T G T C G C C A C T T T A C T C A T A A A T G G	cohl_al2.seq
9386	C C A G C A G A A T C A T C A T A T A A A G T T A C T G T C G C C A C T T T A C T C A T A A A T G G	a909_al2.seq
	A C G A A C A T A A A T T T C T T T T G T C T C A G T T A C A G T T A T T G G C T C A C C A A A T T	Majority
	9460 9470 9480 9490 9500	
9439	A C G A A C A T A A A T T T C T T T T G T C T C A G T T A C A G T T A T T G G C T C A C C A A A T T	cohl_al2.seq
9436	A C G A A C A T A A A T T T C T T T T G T C T C A G T T A C A G T T A T T G G C T C A C C A A A T T	a909_al2.seq
	T A A C A G G G T C A C C A T A C T T T T C C A G T A G T A G G A T C A T A G G T A T A C C A A C C A	Majority
	9510 9520 9530 9540 9550	
9489	T A A C A G G G T C A C C A T A C T T T T C C A G T A G T A G G A T C A T A G G T A T A C C A A C C A	cohl_al2.seq
9486	T A A C A G G G T C A C C A T A C T T T T C C A G T A G T A G G A T C A T A G G T A T A C C A A C C A	a909_al2.seq
	T T A A A A T G C T C T C C T G C T T T A A T C G T C G G A A T C C C A A C T T C T C C T A G A G A	Majority
	9560 9570 9580 9590 9600	
9539	T T A A A A T G C T C T C C T G C T T T A A T C G T C G G A A T C C C A A C T T C T C C T A G A G A	cohl_al2.seq
9536	T T A A A A T G C T C T C C T G C T T T A A T C G T C G G A A T C C C A A C T T C T C C T A G A G A	a909_al2.seq
	T T C T C C A T C T T T T A T A A T T T G A T G A T G A A C T T G C A T A C C T G A A G C T G T C A	Majority
	9610 9620 9630 9640 9650	
9589	T T C T C C A T C T T T T A T A A T T T G A T G A T G A A C T T G C A T A C C T G A A G C T G T C A	cohl_al2.seq
9586	T T C T C C A T C T T T T A T A A T T T G A T G A T G A A C T T G C A T A C C T G A A G C T G T C A	a909_al2.seq
	G G A A A T T A T A A T C A G T T C C G T C A T T A T T T T G A A A A T G G T A A G T T A A C C T A	Majority
	9660 9670 9680 9690 9700	
9639	G G A A A T T A T A A T C A G T T C C G T C A T T A T T T T G A A A A T G G T A A G T T A A C C T A	cohl_al2.seq
9636	G G A A A T T A T A A T C A G T T C C G T C A T T A T T T T G A A A A T G G T A A G T T A A C C T A	a909_al2.seq
	G G A A C T T C T G T A T T A T C C T C T T G A A C A A T T G C A T A A A T G G A G A A T G A A T C	Majority
	9710 9720 9730 9740 9750	
9689	G G A A C T T C T G T A T T A T C C T C T T G A A C A A T T G C A T A A A T G G A G A A T G A A T C	cohl_al2.seq
9686	G G A A C T T C T G T A T T A T C C T C T T G A A C A A T T G C A T A A A T G G A G A A T G A A T C	a909_al2.seq

FIGURE 21N



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	TGTTTTTAAAGCAACATCACTGCTAGTGTCTTCTTAGTTTCTGCAAGTATCTT	Majority
	9760 9770 9780 9790 9800	
9739	TGTTTTTAAAGCAACATCACTGCTAGTGTCTTCTTAGTTTCTGCAAGTATCTT	coh1_al2.seq
9736	TGTTTTTAAAGCAACATCACTGCTAGTGTCTTCTTAGTTTCTGCAAGTATCTT	a909_al2.seq
	TAGATTTTAAATACTTCTGTTTGACCATCATCTTTAAAGTGAACAACCTTTA	Majority
	9810 9820 9830 9840 9850	
9789	TAGATTTTAAATACTTCTGTTTGACCATCATCTTTAAAGTGAACAACCTTTA	coh1_al2.seq
9786	TAGATTTTAAATACTTCTGTTTGACCATCATCTTTAAAGTGAACAACCTTTA	a909_al2.seq
	AGGTTTTTCATCTGAAGCTTCTAATGGCTTATCATAGTTGACCTCTACTTT	Majority
	9860 9870 9880 9890 9900	
9839	AGGTTTTTCATCTGAAGCTTCTAATGGCTTATCATAGTTGACCTCTACTTT	coh1_al2.seq
9836	AGGTTTTTCATCTGAAGCTTCTAATGGCTTATCATAGTTGACCTCTACTTT	a909_al2.seq
	TACTGGGGCTTGGGGTTCTGCTTCTTTTACCATTGTGACTCAATAGTAATGT	Majority
	9910 9920 9930 9940 9950	
9889	TACTGGGGCTTGGGGTTCTGCTTCTTTTACCATTGTGACTCAATAGTAATGT	coh1_al2.seq
9886	TACTGGGGCTTGGGGTTCTGCTTCTTTTACCATTGTGACTCAATAGTAATGT	a909_al2.seq
	CATAGAGTTTGAAGTTTTTGAATTTCACTATCTTGTTTAGCAACTTCTGTC	Majority
	9960 9970 9980 9990 10000	
9939	CATAGAGTTTGAAGTTTTTGAATTTCACTATCTTGTTTAGCAACTTCTGTC	coh1_al2.seq
9936	CATAGAGTTTGAAGTTTTTGAATTTCACTATCTTGTTTAGCAACTTCTGTC	a909_al2.seq
	AATGCTTTTTTCTTATAGTCTTTTAAAGTAGCTGAATTGTCTTTTAATTTC	Majority
	10010 10020 10030 10040 10050	
9989	AATGCTTTTTTCTTATAGTCTTTTAAAGTAGCTGAATTGTCTTTTAATTTC	coh1_al2.seq
9986	AATGCTTTTTTCTTATAGTCTTTTAAAGTAGCTGAATTGTCTTTTAATTTC	a909_al2.seq
	CGTCAACCTTTTAAATCAGCATTTTTAGGAATCTTAGCTTCTTTGGTCAAAG	Majority
	10060 10070 10080 10090 10100	
10039	CGTCAACCTTTTAAATCAGCATTTTTAGGAATCTTAGCTTCTTTGGTCAAAG	coh1_al2.seq
10036	CGTCAACCTTTTAAATCAGCATTTTTAGGAATCTTAGCTTCTTTGGTCAAAG	a909_al2.seq
	TCACTGTTACAGTATAGTCTGCACTCTAAACATCAATGGTTCTTTCACGG	Majority
	10110 10120 10130 10140 10150	
10089	TCACTGTTACAGTATAGTCTGCACTCTAAACATCAATGGTTCTTTCACGG	coh1_al2.seq
10086	TCACTGTTACAGTATAGTCTGCACTCTAAACATCAATGGTTCTTTCACGG	a909_al2.seq
	TAAGCAGCTTCTCAGAAAGATGATGTTTCTGTTACACTAGAAGCAGGAGT	Majority
	10160 10170 10180 10190 10200	
10139	TAAGCAGCTTCTCAGAAAGATGATGTTTCTGTTACACTAGAAGCAGGAGT	coh1_al2.seq
10136	TAAGCAGCTTCTCAGAAAGATGATGTTTCTGTTACACTAGAAGCAGGAGT	a909_al2.seq
	CTGTGGCTTGTCTCTGCTCAACACTTGATTGAGAAGTACATGTTGATGAAG	Majority
	10210 10220 10230 10240 10250	
10189	CTGTGGCTTGTCTCTGCTCAACACTTGATTGAGAAGTACATGTTGATGAAG	coh1_al2.seq
10186	CTGTGGCTTGTCTCTGCTCAACACTTGATTGAGAAGTACATGTTGATGAAG	a909_al2.seq
	TTACCTGGCTAGAAATTTTATTTTCTAAAGTAATCCCCACATCATCTGTC	Majority
	10260 10270 10280 10290 10300	
10239	TTACCTGGCTAGAAATTTTATTTTCTAAAGTAATCCCCACATCATCTGTC	coh1_al2.seq
10236	TTACCTGGCTAGAAATTTTATTTTCTAAAGTAATCCCCACATCATCTGTC	a909_al2.seq
	TTAGTTTCTTCAACTGTTATTGCTGGTAGAATTAATAAATAAGTCGTTAA	Majority
	10310 10320 10330 10340 10350	
10289	TTAGTTTCTTCAACTGTTATTGCTGGTAGAATTAATAAATAAGTCGTTAA	coh1_al2.seq
10286	TTAGTTTCTTCAACTGTTATTGCTGGTAGAATTAATAAATAAGTCGTTAA	a909_al2.seq
	AAAAGTTGTTAGGATCATCAATGACCACATGATAATTTTCCACTCTTTAG	Majority
	10360 10370 10380 10390 10400	
10339	AAAAGTTGTTAGGATCATCAATGACCACATGATAATTTTCCACTCTTTAG	coh1_al2.seq
10336	AAAAGTTGTTAGGATCATCAATGACCACATGATAATTTTCCACTCTTTAG	a909_al2.seq

FIGURE 210

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GGTGTCTTTTCTCTTTTAAATGATTGATTATAAAAAGTTTGACACTTCTTT Majority  
10410 10420 10430 10440 10450  
10389 GGTGTCTTTTCTCTTTTAAATGATTGATTATAAAAAGTTTGACACTTCTTT coh1\_al2.seq  
10386 GGTGTCTTTTCTCTTTTAAATGATTGATTATAAAAAGTTTGACACTTCTTT a909\_al2.seq  
AGCATTCTTTCATCCTCCCTAACCTTAATTGATACTACTAATCTTACCTA Majority  
10460 10470 10480 10490 10500  
10439 AGCATTCTTTCATCCTCCCTAACCTTAATTGATACTACTAATCTTACCTA coh1\_al2.seq  
10436 AGCATTCTTTCATCCTCCCTAACCTTAATTGATACTACTAATCTTACCTA a909\_al2.seq  
GAGGCCATATTCTGAAAGAAATTTTACCTACAATTTGTTCTTCTGAAACA Majority  
10510 10520 10530 10540 10550  
10489 GAGGCCATATTCTGAAAGAAATTTTACCTACAATTTGTTCTTCTGAAACA coh1\_al2.seq  
10485 GAGGCCATATTCTGAAAGAAATTTTACCTACAATTTGTTCTTCTGAAACA a909\_al2.seq  
TCTCCTACAGAAGTATTTTCGAGAATCAATTGAAGTTTTTTCGGTTGTCTCC Majority  
10560 10570 10580 10590 10600  
10539 TCTCCTACAGAAGTATTTTCGAGAATCAATTGAAGTTTTTTCGGTTGTCTCC coh1\_al2.seq  
10535 TCTCCTACAGAAGTATTTTCGAGAATCAATTGAAGTTTTTTCGGTTGTCTCC a909\_al2.seq  
TAATACAAAAATTTTTTTTATCAGGTACTTGAATATGGGTATTTTATATTAC Majority  
10610 10620 10630 10640 10650  
10589 TAATACAAAAATTTTTTTTATCAGGTACTTGAATATGGGTATTTTATATTAC coh1\_al2.seq  
10585 TAATACAAAAATTTTTTTTATCAGGTACTTGAATATGGGTATTTTATATTAC a909\_al2.seq  
TATTACCGAGTGCTTTTATGAATAACATATGGTTCTTTCAACTTATGTTGA Majority  
10660 10670 10680 10690 10700  
10639 TATTACCGAGTGCTTTTATGAATAACATATGGTTCTTTCAACTTATGTTGA coh1\_al2.seq  
10635 TATTACCGAGTGCTTTTATGAATAACATATGGTTCTTTCAACTTATGTTGA a909\_al2.seq  
TTCACGTAAACATCCCTTGAGAATCAATATTAACCCAGTCTCCTGACTC Majority  
10710 10720 10730 10740 10750  
10689 TTCACGTAAACATCCCTTGAGAATCAATATTAACCCAGTCTCCTGACTC coh1\_al2.seq  
10685 TTCACGTAAACATCCCTTGAGAATCAATATTAACCCAGTCTCCTGACTC a909\_al2.seq  
TGCAATAAACCCTTGACTAGGACCTTATTATTGTAGTAAACGCGACAA Majority  
10760 10770 10780 10790 10800  
10739 TGCAATAAACCCTTGACTAGGACCTTATTATTGTAGTAAACGCGACAA coh1\_al2.seq  
10735 TGCAATAAACCCTTGACTAGGACCTTATTATTGTAGTAAACGCGACAA a909\_al2.seq  
CGTCTCCAGTTTTTAAAAATTTGAACCTTTTACTGTAAAGACTACATCACCT Majority  
10810 10820 10830 10840 10850  
10789 CGTCTCCAGTTTTTAAAAATTTGAACCTTTTACTGTAAAGACTACATCACCT coh1\_al2.seq  
10785 CGTCTCCAGTTTTTAAAAATTTGAACCTTTTACTGTAAAGACTACATCACCT a909\_al2.seq  
GCACTTAAAGTCTTATTTCATTGAATGTCCTAGATTCTTAATACAGGCAA Majority  
10860 10870 10880 10890 10900  
10839 GCACTTAAAGTCTTATTTCATTGAATGTCCTAGATTCTTAATACAGGCAA coh1\_al2.seq  
10835 GCACTTAAAGTCTTATTTCATTGAATGTCCTAGATTCTTAATACAGGCAA a909\_al2.seq  
CCATAAAACCGCAATTAAAAATGGCTGTTGAGGCAACCGCCATCAAGATGT Majority  
10910 10920 10930 10940 10950  
10889 CCATAAAACCGCAATTAAAAATGGCTGTTGAGGCAACCGCCATCAAGATGT coh1\_al2.seq  
10885 CCATAAAACCGCAATTAAAAATGGCTGTTGAGGCAACCGCCATCAAGATGT a909\_al2.seq  
ATATGGTATTTTTTAAATGACACTCCAAAAGCGTTTCTGATAAGTTACGGCA Majority  
10960 10970 10980 10990 11000  
10939 ATATGGTATTTTTTAAATGACACTCCAAAAGCGTTTCTGATAAGTTACGGCA coh1\_al2.seq  
10935 ATATGGTATTTTTTAAATGACACTCCAAAAGCGTTTCTGATAAGTTACGGCA a909\_al2.seq  
TCCAGTTCTTGAGATAATTTATCTGAACCTAATCTGTCTTTTCATTGTCTA Majority  
11010 11020 11030 11040 11050  
10989 TCCAGTTCTTGAGATAATTTATCTGAACCTAATCTGTCTTTTCATTGTCTA coh1\_al2.seq  
10985 TCCAGTTCTTGAGATAATTTATCTGAACCTAATCTGTCTTTTCATTGTCTA a909\_al2.seq

FIGURE 21P



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CGCCTCTCTTATCTACTAAATTCTTTACGTTTTCTAAATAGGTATCTGCA Majority
11060 11070 11080 11090 11100
11039 CGCCTCTCTTATCTACTAAATTCTTTACGTTTTCTAAATAGGTATCTGCA coh1_al2.seq
11035 CGCCTCTCTTATCTACTAAATTCTTTACGTTTTCTAAATAGGTATCTGCA a909_al2.seq

GCTAACTGAGCAGTCTCAAAAATACCACTAAGCATTAAAGGAAGCTTCGGC Majority
11110 11120 11130 11140 11150
11089 GCTAACTGAGCAGTCTCAAAAATACCACTAAGCATTAAAGGAAGCTTCGGC coh1_al2.seq
11085 GCTAACTGAGCAGTCTCAAAAATACCACTAAGCATTAAAGGAAGCTTCGGC a909_al2.seq

AATAGAACCAGCTTTGCTAATTTTGAATTTTTTATCATCTAGCGCTTCTT Majority
11160 11170 11180 11190 11200
11139 AATAGAACCAGCTTTGCTAATTTTGAATTTTTTATCATCTAGCGCTTCTT coh1_al2.seq
11135 AATAGAACCAGCTTTGCTAATTTTGAATTTTTTATCATCTAGCGCTTCTT a909_al2.seq

TAAGTTGCTGAATCTCTTTCTCTTGTGTTTTTCAATAAGAAGTTGCTGTTCT Majority
11210 11220 11230 11240 11250
11189 TAAGTTGCTGAATCTCTTTCTCTTGTGTTTTTCAATAAGAAGTTGCTGTTCT coh1_al2.seq
11185 TAAGTTGCTGAATCTCTTTCTCTTGTGTTTTTCAATAAGAAGTTGCTGTTCT a909_al2.seq

AACATAATTTCTAGCAAGTCTTTTCTTTTTTAAATTTTTTAAATCTTCCAT Majority
11260 11270 11280 11290 11300
11239 AACATAATTTCTAGCAAGTCTTTTCTTTTTTAAATTTTTTAAATCTTCCAT coh1_al2.seq
11235 AACATAATTTCTAGCAAGTCTTTTCTTTTTTAAATTTTTTAAATCTTCCAT a909_al2.seq

CGCGATTACTTCCCTTAACTGAACCTTAAATTATCGTTTAGATATTATATC Majority
11310 11320 11330 11340 11350
11289 CGCGATTACTTCCCTTAACTGAACCTTAAATTATCGTTTAGATATTATATC coh1_al2.seq
11285 CGCGATTACTTCCCTTAACTGAACCTTAAATTATCGTTTAGATATTATATC a909_al2.seq

AAAGTTCTAACCCTTTAAACTCATTTTTTGTCTGTTGTTTTTCTCAAAAAA Majority
11360 11370 11380 11390 11400
11339 AAAGTTCTAACCCTTTAAACTCATTTTTTGTCTGTTGTTTTTCTCAAAAAA coh1_al2.seq
11335 AAAGTTCTAACCCTTTAAACTCATTTTTTGTCTGTTGTTTTTCTCAAAAAA a909_al2.seq

AGTCTATGCTAAATTAACATTTTTTGATAATTTTTTGA AAAAATCTCATCGA Majority
11410 11420 11430 11440 11450
11389 AGTCTATGCTAAATTAACATTTTTTGATAATTTTTTGA AAAAATCTCATCGA coh1_al2.seq
11385 AGTCTATGCTAAATTAACATTTTTTGATAATTTTTTGA AAAAATCTCATCGA a909_al2.seq

AGTCATTTTTCTTTTTTGAAGCTCGAATTCTAGGCATTAAAAAGCCATATA Majority
11460 11470 11480 11490 11500
11439 AGTCATTTTTCTTTTTTGAAGCTCGAATTCTAGGCATTAAAAAGCCATATA coh1_al2.seq
11435 AGTCATTTTTCTTTTTTGAAGCTCGAATTCTAGGCATTAAAAAGCCATATA a909_al2.seq

TCAAAATTGATATATGGCTTTTTTTTATTATTAAACAAAAGCAATCAATAG Majority
11510 11520 11530 11540 11550
11489 TCAAAATTGATATATGGCTTTTTTTTATTATTAAACAAAAGCAATCAATAG coh1_al2.seq
11476 TCAAAATTGATATATGGCTTTTTTTTATTATTAAACAAAAGCAATCAATAG a909_al2.seq

GACAATAGCGTCAATTTAGTGACATAATCTATTACAGATTAAAGTTCTTTT Majority
11560 11570 11580 11590 11600
11539 GACAATAGCGTCAATTTAGTGACATAATCTATTACAGATTAAAGTTCTTTT coh1_al2.seq
11492 GACAATAGCGTCAATTTAGTGACATAATCTATTACAGATTAAAGTTCTTTT a909_al2.seq

TGAATAATATAATCCAACCTTTTCAACTGTTTTTTTCCCATGTGAAATGTTTC Majority
11610 11620 11630 11640 11650
11589 TGAATAATATAATCCAACCTTTTCAACTGTTTTTTTCCCATGTGAAATGTTTC coh1_al2.seq
11542 TGAATAATATAATCCAACCTTTTCAACTGTTTTTTTCCCATGTGAAATGTTTC a909_al2.seq

TTTAATTCTTTTAGCAATATTCTGTTGTAGTTTCTCTCTTAATGCCTTAT Majority
11660 11670 11680 11690 11700
11639 TTTAATTCTTTTAGCAATATTCTGTTGTAGTTTCTCTCTTAATGCCTTAT coh1_al2.seq
11592 TTTAATTCTTTTAGCAATATTCTGTTGTAGTTTCTCTCTTAATGCCTTAT a909_al2.seq
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FIGURE 21Q

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CTTTTACTAATAAATAAGAGATTTCATGGAGTGACTGAGTATTTTCTTCC Majority
11710 11720 11730 11740 11750
11689 CTTTACTAATAAATCAAGAGATTTCATGGAGTGACTGAGTATTTTCTTCC coh1_al2.seq
11642 CTTTACTAATAAATCAAGAGATTTCATGGAGTGACTGAGTATTTTCTTCC a909_al2.seq

ATGATGATTCTTAACCTCAGGGCTATCAATAACTTCAACTGTTCCACCGCG Majority
11760 11770 11780 11790 11800
11739 ATGATGATTCTTAACCTCAGGGCTATCAATAACTTCAACTGTTCCACCGCG coh1_al2.seq
11692 ATGATGATTCTTAACCTCAGGGCTATCAATAACTTCAACTGTTCCACCGCG a909_al2.seq

ATCTGTTGCAATAATAGCACTCGAAAGTAGACCAGCTTCTAAAATAGAGG Majority
11810 11820 11830 11840 11850
11789 ATCTGTTGCAATAATAGCACTCGAAAGTAGACCAGCTTCTAAAATAGAGG coh1_al2.seq
11742 ATCTGTTGCAATAATAGCACTCGAAAGTAGACCAGCTTCTAAAATAGAGG a909_al2.seq

TTGGTAATCCCTCTGGATACATTGATGGGTAAACAAAGATATCTGTCTGT Majority
11860 11870 11880 11890 11900
11839 TTGGTAATCCCTCTGGATACATTGATGGGTAAACAAAGATATCTGTCTGT coh1_al2.seq
11792 TTGGTAATCCCTCTGGATACATTGATGGGTAAACAAAGATATCTGTCTGT a909_al2.seq

GCCATTAAAGACATAGTCTGTTCAAAGTTTAAATTTCCCCAAAAAGTTAAT Majority
11910 11920 11930 11940 11950
11889 GCCATTAAAGACATAGTCTGTTCAAAGTTTAAATTTCCCCAAAAAGTTAAT coh1_al2.seq
11842 GCCATTAAAGACATAGTCTGTTCAAAGTTTAAATTTCCCCAAAAAGTTAAT a909_al2.seq

CTGTTTGGACTGATATTTCTCTTTCAAATGTGCTAATTCAGGTCCGTCTC Majority
11960 11970 11980 11990 12000
11939 CTGTTTGGACTGATATTTCTCTTTCAAATGTGCTAATTCAGGTCCGTCTC coh1_al2.seq
11892 CTGTTTGGACTGATATTTCTCTTTCAAATGTGCTAATTCAGGTCCGTCTC a909_al2.seq

CTGCAATCTGTAAATAAACATTTTTCAGAGTACTGTGACATCGAAAAATGCT Majority
12010 12020 12030 12040 12050
11989 CTGCAATCTGTAAATAAACATTTTTCAGAGTACTGTGACATCGAAAAATGCT coh1_al2.seq
11942 CTGCAATCTGTAAATAAACATTTTTCAGAGTACTGTGACATCGAAAAATGCT a909_al2.seq

TCTAAGAGCAATTCAATGCCTTTTTTCTTTAATAATTCTACCAGCATAAGT Majority
12060 12070 12080 12090 12100
12039 TCTAAGAGCAATTCAATGCCTTTTTTCTTTAATAATTCTACCAGCATAAGT coh1_al2.seq
11992 TCTAAGAGCAATTCAATGCCTTTTTTCTTTAATAATTCTACCAGCATAAGT a909_al2.seq

GATGAAAAATATCATCAGCAGATTTTTTCAAGGTAAGCCGTGTCAGCAAAAT Majority
12110 12120 12130 12140 12150
12089 GATGAAAAATATCATCAGCAGATTTTTTCAAGGTAAGCCGTGTCAGCAAAAT coh1_al2.seq
12042 GATGAAAAATATCATCAGCAGATTTTTTCAAGGTAAGCCGTGTCAGCAAAAT a909_al2.seq

CAGAGCCTAGACTTTTCAGATACCGAATTATAAATAACTCCTTTAGCTTCT Majority
12160 12170 12180 12190 12200
12139 CAGAGCCTAGACTTTTCAGATACCGAATTATAAATAACTCCTTTAGCTTCT coh1_al2.seq
12092 CAGAGCCTAGACTTTTCAGATACCGAATTATAAATAACTCCTTTAGCTTCT a909_al2.seq

ATATTAAAAATGTTTTAACCATTCAACGGCTTCTCTTGGATACCGCATAAAA Majority
12210 12220 12230 12240 12250
12189 ATATTAAAAATGTTTTAACCATTCAACGGCTTCTCTTGGATACCGCATAAAA coh1_al2.seq
12142 ATATTAAAAATGTTTTAACCATTCAACGGCTTCTCTTGGATACCGCATAAAA a909_al2.seq

ATCTGGACGATAGTGCTTAACACGGCGCTGTGAGAAGATGTTTCATAGATAG Majority
12260 12270 12280 12290 12300
12239 ATCTGGACGATAGTGCTTAACACGGCGCTGTGAGAAGATGTTTCATAGATAG coh1_al2.seq
12192 ATCTGGACGATAGTGCTTAACACGGCGCTGTGAGAAGATGTTTCATAGATAG a909_al2.seq

CTCCAAAGAAATCTAAAAAACGATTATTGACAGAAAAATGACTTGACCCA Majority
12310 12320 12330 12340 12350
12289 CTCCAAAGAAATCTAAAAAACGATTATTGACAGAAAAATGACTTGACCCA coh1_al2.seq
12242 CTCCAAAGAAATCTAAAAAACGATTATTGACAGAAAAATGACTTGACCCA a909_al2.seq

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FIGURE 21R



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	TGGTCTAAAAACAATACTAGGTAACCGGTGCTTCTTTGCAAAAGATAGCCC	Majority
	12360 12370 12380 12390 12400	
12339	TGGTCTAAAAACAATACTAGGTAACCGGTGCTTCTTTGCAAAAGATAGCCC	cohl_al2.seq
12292	TGGTCTAAAAACAATACTAGGTAACCGGTGCTTCTTTGCAAAAGATAGCCC	a909_al2.seq
	TTCTAGCGTTGTTAACTGAAAACGTGTATTACAAATCACAAAATCAATAT	Majority
	12410 12420 12430 12440 12450	
12389	TTCTAGCGTTGTTAACTGAAAACGTGTATTACAAATCACAAAATCAATAT	cohl_al2.seq
12342	TTCTAGCGTTGTTAACTGAAAACGTGTATTACAAATCACAAAATCAATAT	a909_al2.seq
	TTTCATCTGAAACATGTTTTCATCAGCGTGTGTTGTTTCTCGATTTTTTGTTA	Majority
	12460 12470 12480 12490 12500	
12439	TTTCATCTGAAACATGTTTTCATCAGCGTGTGTTGTTTCTCGATTTTTTGTTA	cohl_al2.seq
12392	TTTCATCTGAAACATGTTTTCATCAGCGTGTGTTGTTTCTCGATTTTTTGTTA	a909_al2.seq
	ATAATAGGATAGCGCTGCTTGACAATGTTTTTGGTCGGTAAACGGTAAAT	Majority
	12510 12520 12530 12540 12550	
12489	ATAATAGGATAGCGCTGCTTGACAATGTTTTTGGTCGGTAAACGGTAAAT	cohl_al2.seq
12442	ATAATAGGATAGCGCTGCTTGACAATGTTTTTGGTCGGTAAACGGTAAAT	a909_al2.seq
	TTTTCTACCCTTGCTTTCATCTATAATCGGTAAATCATCATGATTAGTTG	Majority
	12560 12570 12580 12590 12600	
12539	TTTTCTACCCTTGCTTTCATCTATAATCGGTAAATCATCATGATTAGTTG	cohl_al2.seq
12492	TTTTCTACCCTTGCTTTCATCTATAATCGGTAAATCATCATGATTAGTTG	a909_al2.seq
	TTACAATAACAACACGGTAGCCACGCTTAACCAAATCTGCTGTCATTTTA	Majority
	12610 12620 12630 12640 12650	
12589	TTACAATAACAACACGGTAGCCACGCTTAACCAAATCTGCTGTCATTTTA	cohl_al2.seq
12542	TTACAATAACAACACGGTAGCCACGCTTAACCAAATCTGCTGTCATTTTA	a909_al2.seq
	TCTGTATAACGTTCAATACCTCCGAGGAAGGGTAGATAAATATCCTGAGAA	Majority
	12660 12670 12680 12690 12700	
12639	TCTGTATAACGTTCAATACCTCCGAGGAAGGGTAGATAAATATCCTGAGAA	cohl_al2.seq
12592	TCTGTATAACGTTCAATACCTCCGAGGAAGGGTAGATAAATATCCTGAGAA	a909_al2.seq
	AACAGCAACTGTTTTTACCTTATTTTCCATATTTATCCACTTTTCATCAAT	Majority
	12710 12720 12730 12740 12750	
12689	AACAGCAACTGTTTTTACCTTATTTTCCATATTTATCCACTTTTCATCAAT	cohl_al2.seq
12642	AACAGCAACTGTTTTTACCTTATTTTCCATATTTATCCACTTTTCATCAAT	a909_al2.seq
	AAGCCATCTTTTAAAGCCTTTAATCATAGCAACTAATTTTTTGGCTCTTTTG	Majority
	12760 12770 12780 12790 12800	
12739	AAGCCATCTTTTAAAGCCTTTAATCATAGCAACTAATTTTTTGGCTCTTTTG	cohl_al2.seq
12692	AAGCCATCTTTTAAAGCCTTTAATCATAGCAACTAATTTTTTGGCTCTTTTG	a909_al2.seq
	CTCTTCTGCTACCAACACTCGAACAATTCATTTEGGCATAAATACTAAAT	Majority
	12810 12820 12830 12840 12850	
12789	CTCTTCTGCTACCAACACTCGAACAATTCATTTEGGCATAAATACTAAAT	cohl_al2.seq
12742	CTCTTCTGCTACCAACACTCGAACAATTCATTTEGGCATAAATACTAAAT	a909_al2.seq
	ATTTGFGCCGCTTCTTCTTACCATAATTTTTTTTATAATATAAATCGGCATTG	Majority
	12860 12870 12880 12890 12900	
12839	ATTTGFGCCGCTTCTTCTTACCATAATTTTTTTTATAATATAAATCGGCATTG	cohl_al2.seq
12792	ATTTGFGCCGCTTCTTCTTACCATAATTTTTTTTATAATATAAATCGGCATTG	a909_al2.seq
	CGTATCATGTAATATTTTTCGAAATGGTGAATGATTCAATACATGAAAAAC	Majority
	12910 12920 12930 12940 12950	
12889	CGTATCATGTAATATTTTTCGAAATGGTGAATGATTCAATACATGAAAAAC	cohl_al2.seq
12842	CGTATCATGTAATATTTTTCGAAATGGTGAATGATTCAATACATGAAAAAC	a909_al2.seq
	ATGGCCAAATTTTTTAACTCGTGAAGAGTGTCCAATTTTCGTGTAAAAGAC	Majority
	12960 12970 12980 12990 13000	
12939	ATGGCCAAATTTTTTAACTCGTGAAGAGTGTCCAATTTTCGTGTAAAAGAC	cohl_al2.seq
12892	ATGGCCAAATTTTTTAACTCGTGAAGAGTGTCCAATTTTCGTGTAAAAGAC	a909_al2.seq

FIGURE 21S

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C A A T A A A A T T A A C C T G A T A A A C T C T T A T A T C C C A T C T C T G A C A G A C G A T A A Majority
13010      13020      13030      13040      13050
12989 C A A T A A A A T T A A C C T G A T A A A G T C T T A T A T C C C A T C T C T G A C A G A C G A T A A coh1_al2.seq
12942 C A A T A A A A T T A A C C T G A T A A A G T C T T A T A T C C C A T C T C T G A C A G A C G A T A A a909_al2.seq

T T C A T T T C A G A G T C A A C A A A A T C A A T A A A C A T C T C T T C T G C A A A G C C A G A Majority
13060      13070      13080      13090      13100
13039 T T C A T T T C A G A G T C A A C A A A A T C A A T A A A C A T C T C T T C T G C A A A G C C A G A coh1_al2.seq
12992 T T C A T T T C A G A G T C A A C A A A A T C A A T A A A C A T C T C T T C T G C A A A G C C A G A a909_al2.seq

T G T T T C T T C A A A A A C G C T C G T T T T C A T T A A A G C A G C C G A A G T A A T A C A C T Majority
13110      13120      13130      13140      13150
13089 T G T T T C T T C A A A A A C G C T C G T T T T C A T T A A A G C A G C C G A A G T A A T A C A C T coh1_al2.seq
13042 T G T T T C T T C A A A A A C G C T C G T T T T C A T T A A A G C A G C C G A A G T A A T A C A C T a909_al2.seq

C T T C A A T T T C T T T A T A G T C A A A T T C T T G C A T C A C T A A A T C T T C A C G G T T C Majority
13160      13170      13180      13190      13200
13139 C T T C A A T T T C T T T A T A G T C A A A T T C T T G C A T C A C T A A A T C T T C A C G G T T C coh1_al2.seq
13092 C T T C A A T T T C T T T A T A G T C A A A T T C T T G C A T C A C T A A A T C T T C A C G G T T C a909_al2.seq

A T A T C T T G A T A C A A A C A A G A T A A C A T A C C G A C C T T A G G T A A A T G A A G G T A Majority
13210      13220      13230      13240      13250
13189 A T A T C T T G A T A C A A A C A A G A T A A C A T A C C G A C C T T A G G T A A A A G A A G G T A coh1_al2.seq
13142 A T A T C T T G A T A C A A A C A A G A T A A C A T A C C G A C C T T A G G T A A A T G A A G G T A a909_al2.seq

A T T T T C A T A A T T A T C T A T C A A A T C A C C T A G G A C A A C C G A A T C T T G A T C T A Majority
13260      13270      13280      13290      13300
13239 A T T T T C A T A A T T A T C T A T C A A A T C A C C T A G G A C A A C C G A A T C T T G A T C T A coh1_al2.seq
13192 A T T T T C A T A A T T A T C T A T C A A A T C A C C T A G G A C A A C C G A A T C T T G A T C T A a909_al2.seq

A A G T C A A G A A C C A A T C A A A T T C T T G T G C T A C T G C A A A T T G A C C G A T A C A G Majority
13310      13320      13330      13340      13350
13289 A A G T C A A G A A C C A A T C A A A T T C T T G T G C T A C T G C A A A T T G A C C G A T A C A G coh1_al2.seq
13242 A A G T C A A G A A C C A A T C A A A T T C T T G T G C T A C T G C A A A T T G A C C A A T A C A G a909_al2.seq

T T C A A A G C A T A T G C A A T T C C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G Majority
13360      13370      13380      13390      13400
13339 T T C A A A G C A T A T G C A A T T C C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G coh1_al2.seq
13292 T T C A A A G C A T A T G C A A T T C C T T T A T T T T C T G T T A A A T A A T C A A C A G T T A G a909_al2.seq

G T G C C C C T C T T C A T T A T A A T C G G C C A C T A A T T G A G A A A T T T C T T C C T T A T Majority
13410      13420      13430      13440      13450
13389 G T G C C C C T C T T C A T T A T A A T C G G C C A C T A A T T G A G A A A T T T C T T C C T T A T coh1_al2.seq
13342 G T G C C C C T C T T C A T T A T A A T C G G C C A C T A A T T G A G A A A T T T C T C C T T A T a909_al2.seq

T T T T C G A G C C A T T A T C T A C G A T G T A G A T A T G G C T T A C T T G A G G A T A A A T T Majority
13460      13470      13480      13490      13500
13439 T T T T C G A G C C A T T A T C T A C G A T G T A G A T A T G G C T T A C T T G A G G A T A A A T T coh1_al2.seq
13392 T T T T C G A G C C A T T A T C T A C G A T G T A G A T A T G G C T T A C T T G A G G A T A A A T T a909_al2.seq

G C T C G A A T G T T C T G A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T Majority
13510      13520      13530      13540      13550
13489 G C T C G A A T G T T C T G A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T coh1_al2.seq
13442 G C T C G A A T G T T C T G A T C T A A G C G T T C A A T A T T G G G G T T A A A G G T G A C A A T a909_al2.seq

A C C C G C T A A A T A T T T C A T G T T C T A T G C T C T T T T C T A A A A T C T C T A A A T A A Majority
13560      13570      13580      13590      13600
13539 A C C C G C T A A A T A T T T C A T G T T C T A T G C T C T T T T C T A A A A T C T C T A A A T A A coh1_al2.seq
13492 A C C C G C T A A A T A T T T C A T G T T C T A T G C T C T T T T C T A A A A T C T C T A A A T A A a909_al2.seq

C T G A A T G A C T G G T G C T T T G G T T A T A A A A C G A T A C C G A C A T A G A T A G T T A Majority
13610      13620      13630      13640      13650
13589 C T G A A T G A C T G G T G C T T T G G T T A T A A A A C G A T A C C G A C A T A G A T A G T T A coh1_al2.seq
13542 C T G A A T G A C T G G T G C T T T G G T T A T A A A A C G A T A C C G A C A T A G A T A G T T A a909_al2.seq

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FIGURE 21T



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	CTGCTA	CTAA	AACTTTGA	ATGACATA	ATTTACCA	ATGATACTGAC	ATTTGA	Majority
	13660	13670	13680	13690	13700			
13639	CTGCTA	CTAA	AACTTTGA	ATGACATA	ATTTACCA	ATGATACTGAC	ATTTGA	coh1_al2.seq
13592	CTGCTA	CTAA	AACTTTGA	ATGACATA	ATTTACCA	ATGATACTGAC	ATTTGA	a909_al2.seq
	GTATTG	ATATAA	TAGAGT	ACAGCT	CCACTA	AGAGTAGC	AGCAATTAA	Majority
	13710	13720	13730	13740	13750			
13689	GTATTG	ATATAA	TAGAGT	ACAGCT	CCACTA	AGAGTAGC	AGCAATTAA	coh1_al2.seq
13642	GTATTG	ATATAA	TAGAGT	ACAGCT	CCACTA	AGAGTAGC	AGCAATTAA	a909_al2.seq
	GCGCAG	CATTCC	TCTTGT	TAAATT	CTTTAA	AAAGTAA	ATACATCT	Majority
	13760	13770	13780	13790	13800			
13739	GCGCAG	CATTCC	TCTTGT	TAAATT	CTTTAA	AAAGTAA	ATACATCT	coh1_al2.seq
13692	GCGCAG	CATTCC	TCTTGT	TAAATT	CTTTAA	AAAGTAA	ATACATCT	a909_al2.seq
	AGATAG	CTTGAT	ATAGGG	AGACAATA	AAATTC	AGTAATAA	CTGTAGAG	Majority
	13810	13820	13830	13840	13850			
13789	AGATAG	CTTGAT	ATAGGG	AGACAATA	AAATTC	AGTAATAA	CTGTAGAG	coh1_al2.seq
13742	AGATAG	CTTGAT	ATAGGG	AGACAATA	AAATTC	AGTAATAA	CTGTAGAG	a909_al2.seq
	ATAGCT	CCCCAT	AGCACCT	AAAAAT	TGGTAT	TAAAAAG	TATATTA	Majority
	13860	13870	13880	13890	13900			
13839	ATAGCT	CCCCAT	AGCACCT	AAAAAT	TGGTAT	TAAAAAG	TATATTA	coh1_al2.seq
13792	ATAGCT	CCCCAT	AGCACCT	AAAAAT	TGGTAT	TAAAAAG	TATATTA	a909_al2.seq
	ATTTGCC	ACAAGT	CCAATA	ACTGCC	AGACATT	GTTAGCT	TTTTGT	Majority
	13910	13920	13930	13940	13950			
13889	ATTTGCC	ACAAGT	CCAATA	ACTGCC	AGACATT	GTTAGCT	TTTTGT	coh1_al2.seq
13842	ATTTGCC	ACAAGT	CCAATA	ACTGCC	AGACATT	GTTAGCT	TTTTGT	a909_al2.seq
	TTGAAG	CCAGT	AGATACT	GTTGCC	CTAAAG	CGTTACC	ATAAGAA	Majority
	13960	13970	13980	13990	14000			
13939	TTGAAG	CCAGT	AGATACT	GTTGCC	CTAAAG	CGTTACC	ATAAGAA	coh1_al2.seq
13892	TTGAAG	CCAGT	AGATACT	GTTGCC	CTAAAG	CGTTACC	ATAAGAA	a909_al2.seq
	ATGATC	ATCAAA						Majority
	14010							
13989	ATGATC	ATCAAA						coh1_al2.seq
13942	ATGATC	ATCAAA						a909_al2.seq

Decoration 'Decoration #1': Shade (with solid black) residues that differ from the Consensus.

FIGURE 21U

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M K L S K K L L F S A A V L T M V A G S T V E P V A Q F A T G M S I V R A A E V S Q E R P A K T T V Majority										
10 20 30 40 50										
1	M K L S K K L L F S A A V L T M V A G S T V E P V A Q F A T G M S I V R A A E V S Q E R P A K T T V									sag645_2603.pep
1	M K L S K K L L F S A A V L T M V A G S T V E P V A Q F A T G M S I V R A A E V S Q E R P A K T T V									sag645_a909.pep
1	M K L S K K L L F S A A V L T M V A G S T V E P V A Q F A T G M S I V R A A E V S Q E R P A K T T V									sag645_cjb111.pep
1	M K L S K K L L F S A A V L T M V A G S T V E P V A Q F A T G M S I V R A A E V S Q E R P A K T T V									sag645_coh1.pep
1	M K L S K K L L F S A A V L T M V A G S T V E P V A Q F A T G M S I V R A A E V S Q E R P A K T T V									sag645_nem316.pep
N I Y K L Q A D S Y K S E I T S N G G I E N K D G E V I S N Y A K L G D N V K G L Q G V Q F K R Y K Majority										
60 70 80 90 100										
51	N I Y K L Q A D S Y K S E I T S N G G I E N K D G E V I S N Y A K L G D N V K G L Q G V Q F K R Y K									sag645_2603.pep
51	N I Y K L Q A D S Y K S E I T S N G G I E N K D G E V I S N Y A K L G D N V K G L Q G V Q F K R Y K									sag645_a909.pep
51	N I Y K L Q A D S Y K S E I T S N G G I E N K D G E V I S N Y A K L G D N V K G L Q G V Q F K R Y K									sag645_cjb111.pep
51	N I Y K L Q A D S Y K S E I T S N G G I E N K D G E V I S N Y A K L G D N V K G L Q G V Q F K R Y K									sag645_coh1.pep
51	N I Y K L Q A D S Y K S E I T S N G G I E N K D G E V I S N Y A K L G D N V K G L Q G V Q F K R Y K									sag645_nem316.pep
V K T D I S V D E L K K L T T V E A A D A K V G T I L E E G V S L P Q K T N A Q G L V V D A L D S K Majority										
110 120 130 140 150										
01	V K T D I S V D E L K K L T T V E A A D A K V G T I L E E G V S L P Q K T N A Q G L V V D A L D S K									sag645_2603.pep
01	V K T D I S V D E L K K L T T V E A A D A K V G T I L E E G V S L P Q K T N A Q G L V V D A L D S K									sag645_a909.pep
01	V K T D I S V D E L K K L T T V E A A D A K V G T I L E E G V S L P Q K T N A Q G L V V D A L D S K									sag645_cjb111.pep
01	V K T D I S V D E L K K L T T V E A A D A K V G T I L E E G V S L P Q K T N A Q G L V V D A L D S K									sag645_coh1.pep
01	V K T D I S V D E L K K L T T V E A A D A K V G T I L E E G V S L P Q K T N A Q G L V V D A L D S K									sag645_nem316.pep
S N V R Y L Y V E D L K N S P S N I T K A Y A V P F V L E L P V A N S T G T G F L S E I N I Y P K N Majority										
160 170 180 190 200										
51	S N V R Y L Y V E D L K N S P S N I T K A Y A V P F V L E L P V A N S T G T G F L S E I N I Y P K N									sag645_2603.pep
51	S N V R Y L Y V E D L K N S P S N I T K A Y A V P F V L E L P V A N S T G T G F L S E I N I Y P K N									sag645_a909.pep
51	S N V R Y L Y V E D L K N S P S N I T K A Y A V P F V L E L P V A N S T G T G F L S E I N I Y P K N									sag645_cjb111.pep
51	S N V R Y L Y V E D L K N S P S N I T K A Y A V P F V L E L P V A N S T G T G F L S E I N I Y P K N									sag645_coh1.pep
51	S N V R Y L Y V E D L K N S P S N I T K A Y A V P F V L E L P V A N S T G T G F L S E I N I Y P K N									sag645_nem316.pep
V V T D E P K T D K D V K K L G Q D D A G Y T I G E E F K W F L K S T I P A N L G D Y E K F E I T D Majority										
210 220 230 240 250										
01	V V T D E P K T D K D V K K L G Q D D A G Y T I G E E F K W F L K S T I P A N L G D Y E K F E I T D									sag645_2603.pep
01	V V T D E P K T D K D V K K L G Q D D A G Y T I G E E F K W F L K S T I P A N L G D Y E K F E I T D									sag645_a909.pep
01	V V T D E P K T D K D V K K L G Q D D A G Y T I G E E F K W F L K S T I P A N L G D Y E K F E I T D									sag645_cjb111.pep
01	V V T D E P K T D K D V K K L G Q D D A G Y T I G E E F K W F L K S T I P A N L G D Y E K F E I T D									sag645_coh1.pep
01	V V T D E P K T D K D V K K L G Q D D A G Y T I G E E F K W F L K S T I P A N L G D Y E K F E I T D									sag645_nem316.pep
K F A D G L T Y K S V G K I K I G S K T L N R D E H Y T I D E P T V D N Q N T L K I T F K P E K F K Majority										
260 270 280 290 300										
51	K F A D G L T Y K S V G K I K I G S K T L N R D E H Y T I D E P T V D N Q N T L K I T F K P E K F K									sag645_2603.pep
51	K F A D G L T Y K S V G K I K I G S K T L N R D E H Y T I D E P T V D N Q N T L K I T F K P E K F K									sag645_a909.pep
51	K F A D G L T Y K S V G K I K I G S K T L N R D E H Y T I D E P T V D N Q N T L K I T F K P E K F K									sag645_cjb111.pep
51	K F A D G L T Y K S V G K I K I G S K T L N R D E H Y T I D E P T V D N Q N T L K I T F K P E K F K									sag645_coh1.pep
51	K F A D G L T Y K S V G K I K I G S K T L N R D E H Y T I D E P T V D N Q N T L K I T F K P E K F K									sag645_nem316.pep
E I A E L L K G M T L V K N Q D A L D K A T A N T D D A A F L E I P V A S T I N E K A V L G K A I E Majority										
310 320 330 340 350										
11	E I A E L L K G M T L V K N Q D A L D K A T A N T D D A A F L E I P V A S T I N E K A V L G K A I E									sag645_2603.pep
11	E I A E L L K G M T L V K N Q D A L D K A T A N T D D A A F L E I P V A S T I N E K A V L G K A I E									sag645_a909.pep
11	E I A E L L K G M T L V K N Q D A L D K A T A N T D D A A F L E I P V A S T I N E K A V L G K A I E									sag645_cjb111.pep
11	E I A E L L K G M T L V K N Q D A L D K A T A N T D D A A F L E I P V A S T I N E K A V L G K A I E									sag645_coh1.pep
11	E I A E L L K G M T L V K N Q D A L D K A T A N T D D A A F L E I P V A S T I N E K A V L G K A I E									sag645_nem316.pep
N T F E L Q Y D H T P D K A D N P K P S N P P R K P E V H T G G K R F V K K D S T E T Q T L G G A E Majority										
360 370 380 390 400										
11	N T F E L Q Y D H T P D K A D N P K P S N P P R K P E V H T G G K R F V K K D S T E T Q T L G G A E									sag645_2603.pep
11	N T F E L Q Y D H T P D K A D N P K P S N P P R K P E V H T G G K R F V K K D S T E T Q T L G G A E									sag645_a909.pep
11	N T F E L Q Y D H T P D K A D N P K P S N P P R K P E V H T G G K R F V K K D S T E T Q T L G G A E									sag645_cjb111.pep
11	N T F E L Q Y D H T P D K A D N P K P S N P P R K P E V H T G G K R F V K K D S T E T Q T L G G A E									sag645_coh1.pep
11	N T F E L Q Y D H T P D K A D N P K P S N P P R K P E V H T G G K R F V K K D S T E T Q T L G G A E									sag645_nem316.pep

Figure 22



FDLLASDGTAVKWTDALIKANTNKNYIAGEAVTCQPIKLSHTDGTFEIK Majority  
410 420 430 440 450  
101 FDLLASDGTAVKWTDALIKANTNKNYIAGEAVTCQPIKLSHTDGTFEIK sag645\_2603.pep  
101 FDLLASDGTAVKWTDALIKANTNKNYIAGEAVTCQPIKLSHTDGTFEIK sag645\_a909.pep  
101 FDLLASDGTAVKWTDALIKANTNKNYIAGEAVTCQPIKLSHTDGTFEIK sag645\_cjb111.pep  
101 FDLLASDGTAVKWTDALIKANTNKNYIAGEAVTCQPIKLSHTDGTFEIK sag645\_cohl.pep  
101 FDLLASDGTAVKWTDALIKANTNKNYIAGEAVTCQPIKLSHTDGTFEIK sag645\_nem316.pep

GLAYAVDANAEGTAVTYKLKETKAPEGYVIPDKEIEFTVSQTSYNTKPTD Majority  
460 470 480 490 500  
151 GLAYAVDANAEGTAVTYKLKETKAPEGYVIPDKEIEFTVSQTSYNTKPTD sag645\_2603.pep  
151 GLAYAVDANAEGTAVTYKLKETKAPEGYVIPDKEIEFTVSQTSYNTKPTD sag645\_a909.pep  
151 GLAYAVDANAEGTAVTYKLKETKAPEGYVIPDKEIEFTVSQTSYNTKPTD sag645\_cjb111.pep  
151 GLAYAVDANAEGTAVTYKLKETKAPEGYVIPDKEIEFTVSQTSYNTKPTD sag645\_cohl.pep  
151 GLAYAVDANAEGTAVTYKLKETKAPEGYVIPDKEIEFTVSQTSYNTKPTD sag645\_nem316.pep

ITVDSADATPDTIKNNKRPSIPNTGGIGTAIFVAIGA AVMAFAVKGMKRR Majority  
510 520 530 540 550  
01 ITVDSADATPDTIKNNKRPSIPNTGGIGTAIFVAIGA AVMAFAVKGMKRR sag645\_2603.pep  
01 ITVDSADATPDTIKNNKRPSIPNTGGIGTAIFVAIGA AVMAFAVKGMKRR sag645\_a909.pep  
01 ITVDSADATPDTIKNNKRPSIPNTGGIGTAIFVAIGA AVMAFAVKGMKRR sag645\_cjb111.pep  
01 ITVDSADATPDTIKNNKRPSIPNTGGIGTAIFVAIGA AVMAFAVKGMKRR sag645\_cohl.pep  
01 ITVDSADATPDTIKNNKRPSIPNTGGIGTAIFVAIGA AVMAFAVKGMKRR sag645\_nem316.pep

TKDN

Majority

51 TKDN  
51 TKDN  
51 TKDN  
51 TKDN  
51 TKDN

sag645\_2603.pep  
sag645\_a909.pep  
sag645\_cjb111.pep  
sag645\_cohl.pep  
sag645\_nem316.pep

scoration 'Decoration #1': Shade (with solid black) residues that differ from the Consensus.

FIGURE 22A

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M K K R Q K I W R G L S V T L L I L S Q I P F G I L V Q G E T Q D T N Q A L G K V I V K K T G D N A Majority				
10	20	30	40	50
1 M K K R Q K I W R G L S V T L L I L S Q I P F G I L V Q G E T Q D T N Q A L G K V I V K K T G D N A	sag649_2603.pep			
1 M K K R Q K I W R G L S V T L L I L S Q I P F G I L V Q G E T Q D T N Q A L G K V I V K K T G D N A	sag649_coh1.pep			
1 M K K R Q K I W R G L S V T L L I L S Q I P F G I L V Q G E T Q D T N Q A L G K V I V K K T G D N A	sag649_cjb111.pep			
1 M K K R Q K I W R G L S V T L L I L S Q I P F G I L V Q G E T Q D T N Q A L G K V I V K K T G D N A	sag649_18rs21.pep			
1 M K K R Q K I W R G L S V T L L I L S Q I P F G I L V Q G E T Q D T N Q A L G K V I V K K T G D N A	sag649_nem316.pep			
1 M K K R Q K I W R G L S V T L L I L S Q I P F G I L V Q G E T Q D T N Q A L G K V I V K K T G D N A	sag649_cjb111.pep			
T P L G K A T F V L K N D N D K S E T S H E T V E G S G E A T F E N I K P G D Y T L R E E T A P I G Majority				
60	70	80	90	100
51 T P L G K A T F V L K N D N D K S E T S H E T V E G S G E A T F E N I K P G D Y T L R E E T A P I G	sag649_2603.pep			
51 T P L G K A T F V L K N D N D K S E T S H E T V E G S G E A T F E N I K P G D Y T L R E E T A P I G	sag649_coh1.pep			
51 T P L G K A T F V L K N D N D K S E T S H E T V E G S G E A T F E N I K P G D Y T L R E E T A P I G	sag649_cjb111.pep			
51 T P L G K A T F V L K N D N D K S E T S H E T V E G S G E A T F E N I K P G D Y T L R E E T A P I G	sag649_18rs21.pep			
51 T P L G K A T F V L K N D N D K S E T S H E T V E G S G E A T F E N I K P G D Y T L R E E T A P I G	sag649_nem316.pep			
51 T P L G K A T F V L K N D N D K S E T S H E T V E G S G E A T F E N I K P G D Y T L R E E T A P I G	sag649_cjb111.pep			
Y K K T D K T W K V K V A D N G A T I I E G M D A D K A E K R K E V L N A Q Y P K S A I Y E D T K E Majority				
110	120	130	140	150
101 Y K K T D K T W K V K V A D N G A T I I E G M D A D K A E K R K E V L N A Q Y P K S A I Y E D T K E	sag649_2603.pep			
101 Y K K T D K T W K V K V A D N G A T I I E G M D A D K A E K R K E V L N A Q Y P K S A I Y E D T K E	sag649_coh1.pep			
101 Y K K T D K T W K V K V A D N G A T I I E G M D A D K A E K R K E V L N A Q Y P K S A I Y E D T K E	sag649_cjb111.pep			
101 Y K K T D K T W K V K V A D N G A T I I E G M D A D K A E K R K E V L N A Q Y P K S A I Y E D T K E	sag649_18rs21.pep			
101 Y K K T D K T W K V K V A D N G A T I I E G M D A D K A E K R K E V L N A Q Y P K S A I Y E D T K E	sag649_nem316.pep			
101 Y K K T D K T W K V K V A D N G A T I I E G M D A D K A E K R K E V L N A Q Y P K S A I Y E D T K E	sag649_cjb111.pep			
N Y P L V N V E G S K V G E Q Y K A L N P I N G K D G R R E I A E G W L S K K I T G V N D L D K N K Majority				
160	170	180	190	200
151 N Y P L V N V E G S K V G E Q Y K A L N P I N G K D G R R E I A E G W L S K K I T G V N D L D K N K	sag649_2603.pep			
151 N Y P L V N V E G S K V G E Q Y K A L N P I N G K D G R R E I A E G W L S K K I T G V N D L D K N K	sag649_coh1.pep			
151 N Y P L V N V E G S K V G E Q Y K A L N P I N G K D G R R E I A E G W L S K K I T G V N D L D K N K	sag649_cjb111.pep			
151 N Y P L V N V E G S K V G E Q Y K A L N P I N G K D G R R E I A E G W L S K K I T G V N D L D K N K	sag649_18rs21.pep			
151 N Y P L V N V E G S K V G E Q Y K A L N P I N G K D G R R E I A E G W L S K K I T G V N D L D K N K	sag649_nem316.pep			
151 N Y P L V N V E G S K V G E Q Y K A L N P I N G K D G R R E I A E G W L S K K I T G V N D L D K N K	sag649_cjb111.pep			
Y K I E L T V E G K T T V E T K E L N Q P L D V V V L L D N S N S M N N E R A N N S Q R A L K A G E Majority				
210	220	230	240	250
201 Y K I E L T V E G K T T V E T K E L N Q P L D V V V L L D N S N S M N N E R A N N S Q R A L K A G E	sag649_2603.pep			
201 Y K I E L T V E G K T T V E T K E L N Q P L D V V V L L D N S N S M N N E R A N N S Q R A L K A G E	sag649_coh1.pep			
201 Y K I E L T V E G K T T V E T K E L N Q P L D V V V L L D N S N S M N N E R A N N S Q R A L K A G E	sag649_cjb111.pep			
201 Y K I E L T V E G K T T V E T K E L N Q P L D V V V L L D N S N S M N N E R A N N S Q R A L K A G E	sag649_18rs21.pep			
201 Y K I E L T V E G K T T V E T K E L N Q P L D V V V L L D N S N S M N N E R A N N S Q R A L K A G E	sag649_nem316.pep			
201 Y K I E L T V E G K T T V E T K E L N Q P L D V V V L L D N S N S M N N E R A N N S Q R A L K A G E	sag649_cjb111.pep			
A V E K L I D K I T S N K D N R V A L V T Y A S T I F D G T E A T V S K G V A D Q N G K A L N D S V Majority				
260	270	280	290	300
251 A V E K L I D K I T S N K D N R V A L V T Y A S T I F D G T E A T V S K G V A D Q N G K A L N D S V	sag649_2603.pep			
251 A V E K L I D K I T S N K D N R V A L V T Y A S T I F D G T E A T V S K G V A D Q N G K A L N D S V	sag649_coh1.pep			
251 A V E K L I D K I T S N K D N R V A L V T Y A S T I F D G T E A T V S K G V A D Q N G K A L N D S V	sag649_cjb111.pep			
251 A V E K L I D K I T S N K D N R V A L V T Y A S T I F D G T E A T V S K G V A D Q N G K A L N D S V	sag649_18rs21.pep			
251 A V E K L I D K I T S N K D N R V A L V T Y A S T I F D G T E A T V S K G V A D Q N G K A L N D S V	sag649_nem316.pep			
251 A V E K L I D K I T S N K D N R V A L V T Y A S T I F D G T E A T V S K G V A D Q N G K A L N D S V	sag649_cjb111.pep			
S W D Y H K T T F T A T T H N Y S Y L N L T N D A N E V N I L K S R I P K E A E H I N G D R T L Y Q Majority				
310	320	330	340	350
301 S W D Y H K T T F T A T T H N Y S Y L N L T N D A N E V N I L K S R I P K E A E H I N G D R T L Y Q	sag649_2603.pep			
301 S W D Y H K T T F T A T T H N Y S Y L N L T N D A N E V N I L K S R I P K E A E H I N G D R T L Y Q	sag649_coh1.pep			
301 S W D Y H K T T F T A T T H N Y S Y L N L T N D A N E V N I L K S R I P K E A E H I N G D R T L Y Q	sag649_cjb111.pep			
301 S W D Y H K T T F T A T T H N Y S Y L N L T N D A N E V N I L K S R I P K E A E H I N G D R T L Y Q	sag649_18rs21.pep			
301 S W D Y H K T T F T A T T H N Y S Y L N L T N D A N E V N I L K S R I P K E A E H I N G D R T L Y Q	sag649_nem316.pep			
301 S W D Y H K T T F T A T T H N Y S Y L N L T N D A N E V N I L K S R I P K E A E H I N G D R T L Y Q	sag649_cjb111.pep			
F G A T F T Q K A L M K A N E I L E T Q S S N A R K K L I F H V T D G V P T M S Y A I N F N P Y I S Majority				
360	370	380	390	400
351 F G A T F T Q K A L M K A N E I L E T Q S S N A R K K L I F H V T D G V P T M S Y A I N F N P Y I S	sag649_2603.pep			
351 F G A T F T Q K A L M K A N E I L E T Q S S N A R K K L I F H V T D G V P T M S Y A I N F N P Y I S	sag649_coh1.pep			
351 F G A T F T Q K A L M K A N E I L E T Q S S N A R K K L I F H V T D G V P T M S Y A I N F N P Y I S	sag649_cjb111.pep			
351 F G A T F T Q K A L M K A N E I L E T Q S S N A R K K L I F H V T D G V P T M S Y A I N F N P Y I S	sag649_18rs21.pep			
351 F G A T F T Q K A L M K A N E I L E T Q S S N A R K K L I F H V T D G V P T M S Y A I N F N P Y I S	sag649_nem316.pep			
351 F G A T F T Q K A L M K A N E I L E T Q S S N A R K K L I F H V T D G V P T M S Y A I N F N P Y I S	sag649_cjb111.pep			

Figure 23



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TSYQNQFNSFLNKIPDRSGILQEDFIINGDDYQIVKGDGESFKLFSDRKV Majority									
410		420		430		440		450	
401	TSYQNQFNSFLNKIPDRSGILQEDFIINGDDYQIVKGDGESFKLFSDRKV	sag649_2603.pep							
401	TSYQNQFNSFLNKIPDRSGILQEDFIINGDDYQIVKGDGESFKLFSDRKV	sag649_coh1.pep							
401	TSYQNQFNSFLNKIPDRSGILQEDFIINGDDYQIVKGDGESFKLFSDRKV	sag649_cjb111.pep							
401	TSYQNQFNSFLNKIPDRSGILQEDFIINGDDYQIVKGDGESFKLFSDRKV	sag649_18rs21.pep							
401	TSYQNQFNSFLNKIPDRSGILQEDFIINGDDYQIVKGDGESFKLFSDRKV	sag649_nem316.pep							
401	TSYQNQFNSFLNKIPDRSGILQEDFIINGDDYQIVKGDGESFKLFSDRKV	sag649_cjb111.pep							
PVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPPFDPKT Majority									
460		470		480		490		500	
451	PVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPPFDPKT	sag649_2603.pep							
451	PVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPPFDPKT	sag649_coh1.pep							
451	PVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPPFDPKT	sag649_cjb111.pep							
451	PVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPPFDPKT	sag649_18rs21.pep							
451	PVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPPFDPKT	sag649_nem316.pep							
451	PVTGGTTQAAYRVPQNQLSVMSNEGYAINSGYIYLYWRDYNWVYPPFDPKT	sag649_cjb111.pep							
KKVSATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAE Majority									
510		520		530		540		550	
501	KKVSATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAE	sag649_2603.pep							
501	KKVSATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAE	sag649_coh1.pep							
501	KKVSATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAE	sag649_cjb111.pep							
501	KKVSATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAE	sag649_18rs21.pep							
501	KKVSATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAE	sag649_nem316.pep							
501	KKVSATKQIKTHGEPTTLYFNGNIRPKGYDIFTVGIGVNGDPGATPLEAE	sag649_cjb111.pep							
KFMQSISSKTENYTNVDDTNKIYDELNKFYKTIIVEEKHSIVDGNVTDPMG Majority									
560		570		580		590		600	
551	KFMQSISSKTENYTNVDDTNKIYDELNKFYKTIIVEEKHSIVDGNVTDPMG	sag649_2603.pep							
551	KFMQSISSKTENYTNVDDTNKIYDELNKFYKTIIVEEKHSIVDGNVTDPMG	sag649_coh1.pep							
551	KFMQSISSKTENYTNVDDTNKIYDELNKFYKTIIVEEKHSIVDGNVTDPMG	sag649_cjb111.pep							
551	KFMQSISSKTENYTNVDDTNKIYDELNKFYKTIIVEEKHSIVDGNVTDPMG	sag649_18rs21.pep							
551	KFMQSISSKTENYTNVDDTNKIYDELNKFYKTIIVEEKHSIVDGNVTDPMG	sag649_nem316.pep							
551	KFMQSISSKTENYTNVDDTNKIYDELNKFYKTIIVEEKHSIVDGNVTDPMG	sag649_cjb111.pep							
EMIEFQLKNGQSFTHDDYVLVGNDCGSQLKNGVALGGPNSDGGILKDVTVT Majority									
610		620		630		640		650	
601	EMIEFQLKNGQSFTHDDYVLVGNDCGSQLKNGVALGGPNSDGGILKDVTVT	sag649_2603.pep							
601	EMIEFQLKNGQSFTHDDYVLVGNDCGSQLKNGVALGGPNSDGGILKDVTVT	sag649_coh1.pep							
601	EMIEFQLKNGQSFTHDDYVLVGNDCGSQLKNGVALGGPNSDGGILKDVTVT	sag649_cjb111.pep							
601	EMIEFQLKNGQSFTHDDYVLVGNDCGSQLKNGVALGGPNSDGGILKDVTVT	sag649_18rs21.pep							
601	EMIEFQLKNGQSFTHDDYVLVGNDCGSQLKNGVALGGPNSDGGILKDVTVT	sag649_nem316.pep							
601	EMIEFQLKNGQSFTHDDYVLVGNDCGSQLKNGVALGGPNSDGGILKDVTVT	sag649_cjb111.pep							
YDKTSQTIKINHNLGSGQKVVLTYDVRLLKDNYSISNKFYNTNNRTTTLSPK Majority									
660		670		680		690		700	
651	YDKTSQTIKINHNLGSGQKVVLTYDVRLLKDNYSISNKFYNTNNRTTTLSPK	sag649_2603.pep							
651	YDKTSQTIKINHNLGSGQKVVLTYDVRLLKDNYSISNKFYNTNNRTTTLSPK	sag649_coh1.pep							
651	YDKTSQTIKINHNLGSGQKVVLTYDVRLLKDNYSISNKFYNTNNRTTTLSPK	sag649_cjb111.pep							
651	YDKTSQTIKINHNLGSGQKVVLTYDVRLLKDNYSISNKFYNTNNRTTTLSPK	sag649_18rs21.pep							
651	YDKTSQTIKINHNLGSGQKVVLTYDVRLLKDNYSISNKFYNTNNRTTTLSPK	sag649_nem316.pep							
651	YDKTSQTIKINHNLGSGQKVVLTYDVRLLKDNYSISNKFYNTNNRTTTLSPK	sag649_cjb111.pep							
SEKEPNTIRDFPIPKIRDVREFFVLTISNQKKMGEEVEFIKVNKDKHSESL Majority									
710		720		730		740		750	
701	SEKEPNTIRDFPIPKIRDVREFFVLTISNQKKMGEEVEFIKVNKDKHSESL	sag649_2603.pep							
701	SEKEPNTIRDFPIPKIRDVREFFVLTISNQKKMGEEVEFIKVNKDKHSESL	sag649_coh1.pep							
701	SEKEPNTIRDFPIPKIRDVREFFVLTISNQKKMGEEVEFIKVNKDKHSESL	sag649_cjb111.pep							
701	SEKEPNTIRDFPIPKIRDVREFFVLTISNQKKMGEEVEFIKVNKDKHSESL	sag649_18rs21.pep							
701	SEKEPNTIRDFPIPKIRDVREFFVLTISNQKKMGEEVEFIKVNKDKHSESL	sag649_nem316.pep							
701	SEKEPNTIRDFPIPKIRDVREFFVLTISNQKKMGEEVEFIKVNKDKHSESL	sag649_cjb111.pep							
LGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDGNKLYEIS Majority									
760		770		780		790		800	
751	LGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDGNKLYEIS	sag649_2603.pep							
751	LGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDGNKLYEIS	sag649_coh1.pep							
751	LGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDGNKLYEIS	sag649_cjb111.pep							
751	LGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDGNKLYEIS	sag649_18rs21.pep							
751	LGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDGNKLYEIS	sag649_nem316.pep							
751	LGAKFQLQIEKDFSGYKQFVPEGS DVTTKNDGKIYFKALQDGNKLYEIS	sag649_cjb111.pep							

FIGURE 23A

sag649\_2603.pep  
sag649\_coh1.pep  
sag649\_cjb111.pep  
sag649\_18rs21.pep  
sag649\_nen316.pep  
sag649\_cjb111.pep

**FIGURE 23B**



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MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTTDDQ Majority									
10		20		30		40		50	
1	MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTTDDQ	sag1408_2603.pep							
1	MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTTDDQ	sag1408_515.pep							
1	MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTTDDQ	sag1408_18rs21.seq							
1	MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTTDDQ	sag1408_cjb111.pep							
1	MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTTDDQ	sag1408_h36b.pep							
1	MRKYQKFSKILTLSLFCLSQIPLNTNVLGESTVPENGAKGKLVVKKTTDDQ	sag1408_nem316.pep							
NKPLSKATFVLKTTAHPESKIEKVTAELTGEATFDNLIPGDYTLSEETAP Majority									
60		70		80		90		100	
51	NKPLSKATFVLKTTAHPESKIEKVTAELTGEATFDNLIPGDYTLSEETAP	sag1408_2603.pep							
51	NKPLSKATFVLKTTAHPESKIEKVTAELTGEATFDNLIPGDYTLSEETAP	sag1408_515.pep							
51	NKPLSKATFVLKTTAHPESKIEKVTAELTGEATFDNLIPGDYTLSEETAP	sag1408_18rs21.seq							
51	NKPLSKATFVLKTTAHPESKIEKVTAELTGEATFDNLIPGDYTLSEETAP	sag1408_cjb111.pep							
51	NKPLSKATFVLKTTAHPESKIEKVTAELTGEATFDNLIPGDYTLSEETAP	sag1408_h36b.pep							
51	NKPLSKATFVLKTTAHPESKIEKVTAELTGEATFDNLIPGDYTLSEETAP	sag1408_nem316.pep							
EGYKKTNTQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYPPTGIYE Majority									
110		120		130		140		150	
101	EGYKKTNTQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYPPTGIYE	sag1408_2603.pep							
101	EGYKKTNTQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYPPTGIYE	sag1408_515.pep							
101	EGYKKTNTQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYPPTGIYE	sag1408_18rs21.seq							
101	EGYKKTNTQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYPPTGIYE	sag1408_cjb111.pep							
101	EGYKKTNTQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYPPTGIYE	sag1408_h36b.pep							
101	EGYKKTNTQTWQVKVESNGKTTIQNSGDKNSTIGQNQEELDKQYPPTGIYE	sag1408_nem316.pep							
DTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRRISEV Majority									
160		170		180		190		200	
151	DTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRRISEV	sag1408_2603.pep							
151	DTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRRISEV	sag1408_515.pep							
151	DTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRRISEV	sag1408_18rs21.seq							
151	DTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRRISEV	sag1408_cjb111.pep							
151	DTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRRISEV	sag1408_h36b.pep							
151	DTKESYKLEHVKGSPNGKSEAKAVNPYSSEGEHIREIPEGTLSKRRISEV	sag1408_nem316.pep							
GDLAHNKYKIELTVSGKTIKPVVDKQKPLDVVFVLDNSNSMNNNDGPNFQR Majority									
210		220		230		240		250	
201	GDLAHNKYKIELTVSGKTIKPVVDKQKPLDVVFVLDNSNSMNNNDGPNFQR	sag1408_2603.pep							
201	GDLAHNKYKIELTVSGKTIKPVVDKQKPLDVVFVLDNSNSMNNNDGPNFQR	sag1408_515.pep							
201	GDLAHNKYKIELTVSGKTIKPVVDKQKPLDVVFVLDNSNSMNNNDGPNFQR	sag1408_18rs21.seq							
201	GDLAHNKYKIELTVSGKTIKPVVDKQKPLDVVFVLDNSNSMNNNDGPNFQR	sag1408_cjb111.pep							
201	GDLAHNKYKIELTVSGKTIKPVVDKQKPLDVVFVLDNSNSMNNNDGPNFQR	sag1408_h36b.pep							
201	GDLAHNKYKIELTVSGKTIKPVVDKQKPLDVVFVLDNSNSMNNNDGPNFQR	sag1408_nem316.pep							
HNKAKKAAEALGTAVKDILGANSNDRVALVTYGSDFDGRSVDVVKGFKKE Majority									
260		270		280		290		300	
251	HNKAKKAAEALGTAVKDILGANSNDRVALVTYGSDFDGRSVDVVKGFKKE	sag1408_2603.pep							
251	HNKAKKAAEALGTAVKDILGANSNDRVALVTYGSDFDGRSVDVVKGFKKE	sag1408_515.pep							
251	HNKAKKAAEALGTAVKDILGANSNDRVALVTYGSDFDGRSVDVVKGFKKE	sag1408_18rs21.seq							
251	HNKAKKAAEALGTAVKDILGANSNDRVALVTYGSDFDGRSVDVVKGFKKE	sag1408_cjb111.pep							
251	HNKAKKAAEALGTAVKDILGANSNDRVALVTYGSDFDGRSVDVVKGFKKE	sag1408_h36b.pep							
251	HNKAKKAAEALGTAVKDILGANSNDRVALVTYGSDFDGRSVDVVKGFKKE	sag1408_nem316.pep							
DDKYYGLQTKFTIQTENYSHKQLTNNAEEIIRIPTEAPKAKWGSTTNGL Majority									
310		320		330		340		350	
301	DDKYYGLQTKFTIQTENYSHKQLTNNAEEIIRIPTEAPKAKWGSTTNGL	sag1408_2603.pep							
301	DDKYYGLQTKFTIQTENYSHKQLTNNAEEIIRIPTEAPKAKWGSTTNGL	sag1408_515.pep							
301	DDKYYGLQTKFTIQTENYSHKQLTNNAEEIIRIPTEAPKAKWGSTTNGL	sag1408_18rs21.seq							
301	DDKYYGLQTKFTIQTENYSHKQLTNNAEEIIRIPTEAPKAKWGSTTNGL	sag1408_cjb111.pep							
297	DDKYYGLQTKFTIQTENYSHKQLTNNAEEIIRIPTEAPKAKWGSTTNGL	sag1408_h36b.pep							
301	DDKYYGLQTKFTIQTENYSHKQLTNNAEEIIRIPTEAPKAKWGSTTNGL	sag1408_nem316.pep							
TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTDCGVPTR Majority									
360		370		380		390		400	
351	TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTDCGVPTR	sag1408_2603.pep							
351	TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTDCGVPTR	sag1408_515.pep							
351	TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTDCGVPTR	sag1408_18rs21.seq							
351	TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTDCGVPTR	sag1408_cjb111.pep							
346	TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTDCGVPTR	sag1408_h36b.pep							
351	TPEQQKEYYLSKVGETFTMKAFMEADDILSQVNRNSQKIIVHVTDCGVPTR	sag1408_nem316.pep							

Figure 24



S Y A I N N F K L G A S Y E S Q F E Q M K K N G Y L N K S N F L L T D K P E D I K G N G E S Y F L F Majority									
410 420 430 440 450									
401	S	Y	A	I	N	N	F	K	L
401	S	Y	A	I	N	N	F	K	L
401	S	Y	A	I	N	N	F	K	L
401	S	Y	A	I	N	N	F	K	L
396	S	Y	A	I	N	S	F	V	R
401	S	Y	A	I	N	N	F	K	L
P L D S Y Q T Q I I S G N L Q K L H Y L D L N L N Y P K G T I Y R N G P V K E H G T P T K L Y I N S Majority									
460 470 480 490 500									
451	P	L	D	S	Y	Q	T	Q	I
451	P	L	D	S	Y	Q	T	Q	I
451	P	L	D	S	Y	Q	T	Q	I
451	P	L	D	S	Y	Q	T	Q	I
446	P	L	D	S	Y	Q	T	Q	I
451	P	L	D	S	Y	Q	T	Q	I
L K Q K N Y D I F N F G I D I S G F R Q V Y N E D Y K K N Q D G T F Q K L K E E A F E L S D G E I T Majority									
510 520 530 540 550									
501	L	K	Q	K	N	Y	D	I	F
501	L	K	Q	K	N	Y	D	I	F
501	L	K	Q	K	N	Y	D	I	F
501	L	K	Q	K	N	Y	D	I	F
496	L	K	Q	K	N	Y	D	I	F
501	L	K	Q	K	N	Y	D	I	F
E L M R S F S S K P E Y Y T P I V T S A D T S N N E I L S K I Q Q Q F E T I L T K E N S I V N G T I Majority									
560 570 580 590 600									
551	E	L	M	R	S	F	S	S	K
551	E	L	M	R	S	F	S	S	K
551	E	L	M	R	S	F	S	S	K
551	E	L	M	R	S	F	S	S	K
546	E	L	M	R	S	F	S	S	K
551	E	L	M	R	S	F	S	S	K
E D P M G D K I N L Q L G N G Q T L Q P S D Y T L Q G N D G S V M K D G I A T G C P N N D G G I L K Majority									
610 620 630 640 650									
601	E	D	P	M	G	D	K	I	N
601	E	D	P	M	G	D	K	I	N
601	E	D	P	M	G	D	K	I	N
601	E	D	P	M	G	D	K	I	N
596	E	D	P	M	G	D	K	I	N
601	E	D	P	M	G	D	K	I	N
G V K L E Y I G N K L Y V R G L N L G E G Q K V T L T Y D V K L D D S F I S N K F Y D T N G R T T L Majority									
660 670 680 690 700									
651	G	V	K	L	E	Y	I	G	N
651	G	V	K	L	E	Y	I	G	N
651	G	V	K	L	E	Y	I	G	N
651	G	V	K	L	E	Y	I	G	N
646	G	V	K	L	E	Y	I	G	N
651	G	V	K	L	E	Y	I	G	N
N P K S E D P N T L R D F P I P K I R D V R E Y P T I T I K N E K K L G E I E F I K V D K D N N K L Majority									
710 720 730 740 750									
701	N	P	K	S	E	D	P	N	T
701	N	P	K	S	E	D	P	N	T
701	N	P	K	S	E	D	P	N	T
701	N	P	K	S	E	D	P	N	T
696	N	P	K	S	E	D	P	N	T
701	N	P	K	S	E	D	P	N	T
L L K G A T F E L Q E F N E D Y K L Y L P I K N N N S K V V T G E N G K I S Y K D L K D G K Y Q L I Majority									
760 770 780 790 800									
751	L	L	K	G	A	T	F	E	L
751	L	L	K	G	A	T	F	E	L
751	L	L	K	G	A	T	F	E	L
751	L	L	K	G	A	T	F	E	L
746	L	L	K	G	A	T	F	E	L
751	L	L	K	G	A	T	F	E	L

FIGURE 24A



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```
EA V S P E D Y Q K I T N K P I L T F E V V K G S I Q N I I A V N K Q I S E Y H E E G D K H L I T N Majority
      810      820      830      840      850
801 EAVSPEDYQK I T N K P I L T F E V V K G S I Q N I I A V N K Q I S E Y H E E G D K H L I T N sag1408_2603.pep
801 EAVSPEDYQK I T N K P I L T F E V V K G S I Q N I I A V N K Q I S E Y H E E G D K H L I T N sag1408_515.pep
801 EAVSPEDYQK I T N K P I L T F E V V K G S I Q N I I A V N K Q I S E Y H E E G D K H L I T N sag1408_18rs21.seq
801 EAVSPEDYQK I T N K P I L T F E V V K G S I Q N I I A V N K Q I S E Y H E E G D K H L I T N sag1408_cjb111.pep
796 EAVSPEDYQK I T N K P I L T F E V V K G S I Q N I I A V N K Q I S E Y H E E G D K H L I T N sag1408_h36b.pep
801 EAVSPEDYQK I T N K P I L T F E V V K G S I Q N I I A V N K Q I S E Y H E E G D K H L I T N sag1408_nem316.pep
```

```
TH I P P K G I I P M T G G K G I L S F I L I G G A M M S I A G G I Y I W K R Y K K S S D M S I E K Majority
      860      870      880      890      900
851 THIPPKGIIPMTGGKGILS FILIGGAMMSIAGGIYIWKRYKKSSDMSI K K sag1408_2603.pep
851 THIPPKGIIPMTGGKGILS FILIGGAMMSIAGGIYIWKRYKKSSDMSI K K sag1408_515.pep
851 THIPPKGIIPMTGGKGILS FILIGGAMMSIAGGIYIWKRYKKSSDMSI K K sag1408_18rs21.seq
851 THIPPKGIIPMTGGKGILS FILIGGAMMSIAGGIYIWKRYKKSSDMSI K K sag1408_cjb111.pep
846 THIPPKGIIPMTGGKGILS FILIGGAMMSIAGGIYIWKRYKKSSDMSI K K sag1408_h36b.pep
851 THIPPKGIIPMTGGKGILS FILIGGAMMSIAGGIYIWKRYKKSSDMSI K K sag1408_nem316.pep
```

D

Majority

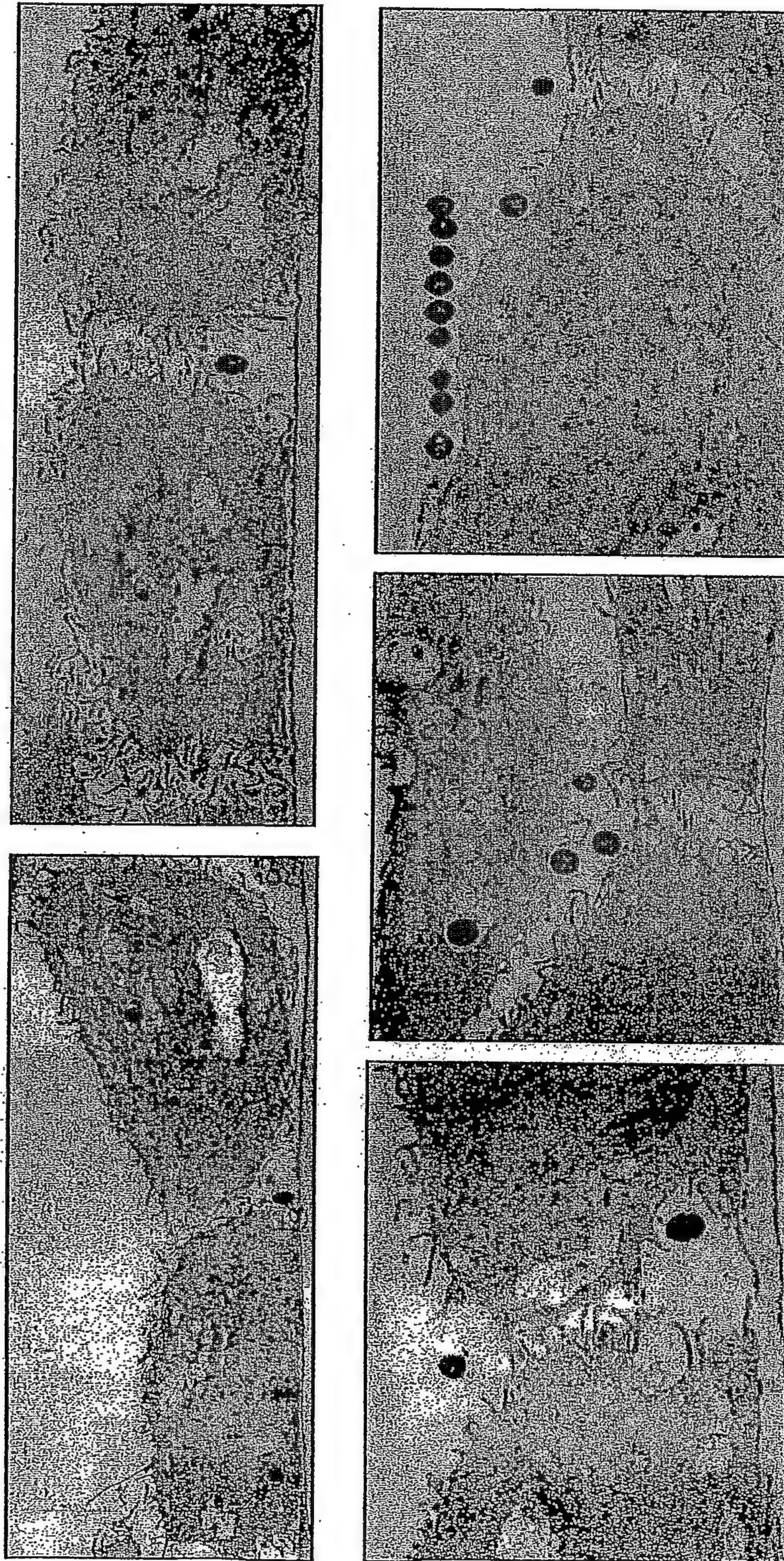
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901 D
901 D
901 D
901 D
896 D
901 D
sag1408_2603.pep
sag1408_515.pep
sag1408_18rs21.seq
sag1408_cjb111.pep
sag1408_h36b.pep
sag1408_nem316.pep
```

Decoration 'Decoration #1': Shade (with solid black) residues that differ from the Consensus.

FIGURE 24B



Figure 25: GBS closely associate with tight junctions and cross the monolayer by a paracellular route



Transmission Electron Microscopy images of GBS infection of ME180 cervical epithelial cells.



Figure 26: GBS infection of ME180 cells  
GBS infection of ME180 cells

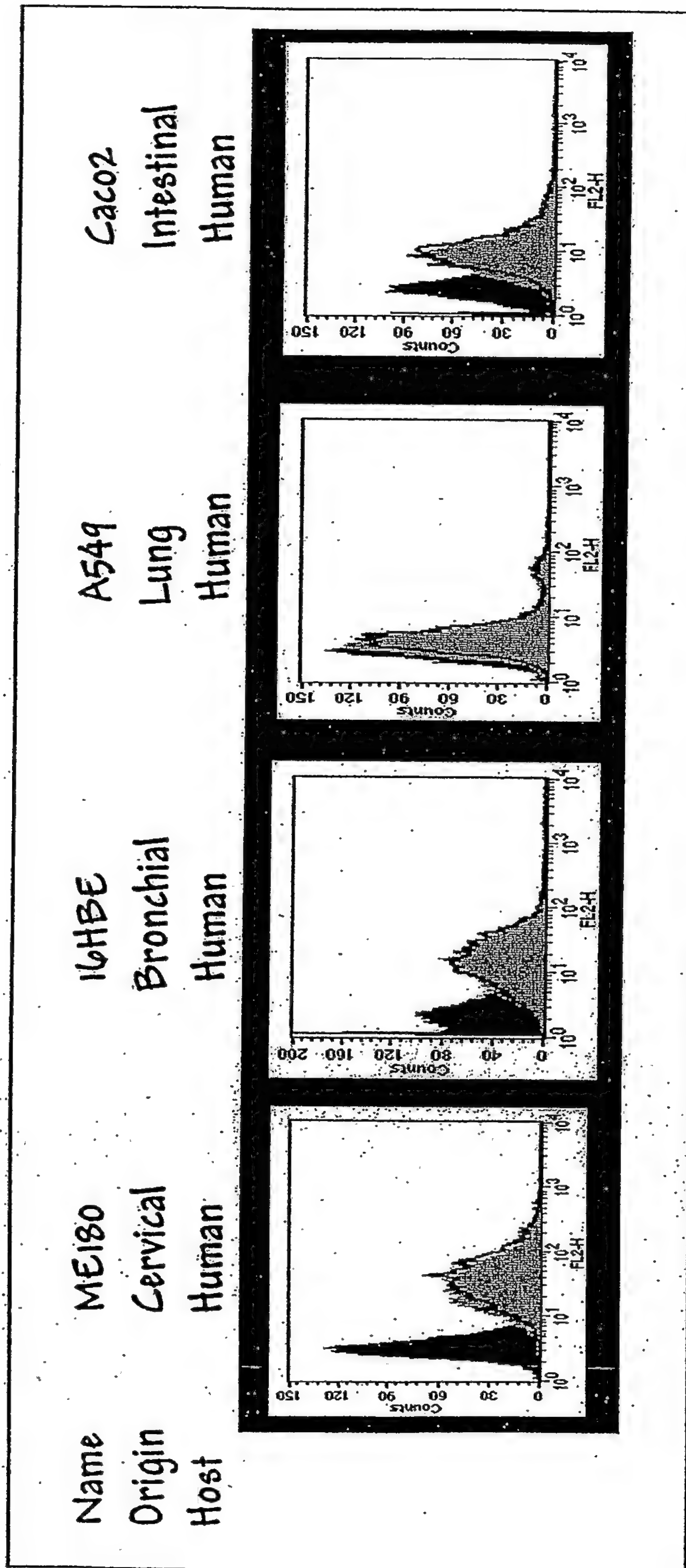


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PCT/US05/27239

PCT/US05/27239/142/487

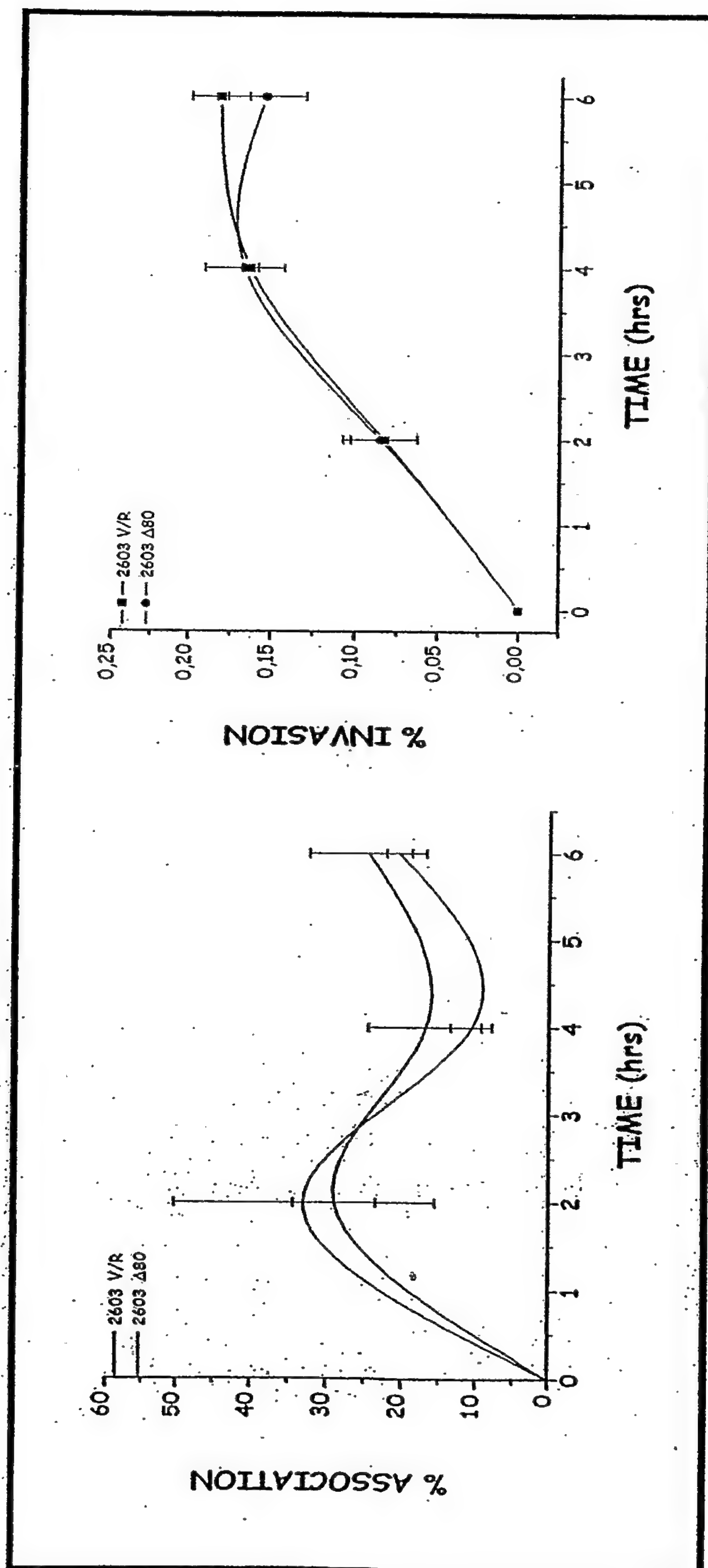
Figure 27





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Figure 28



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Figure 29

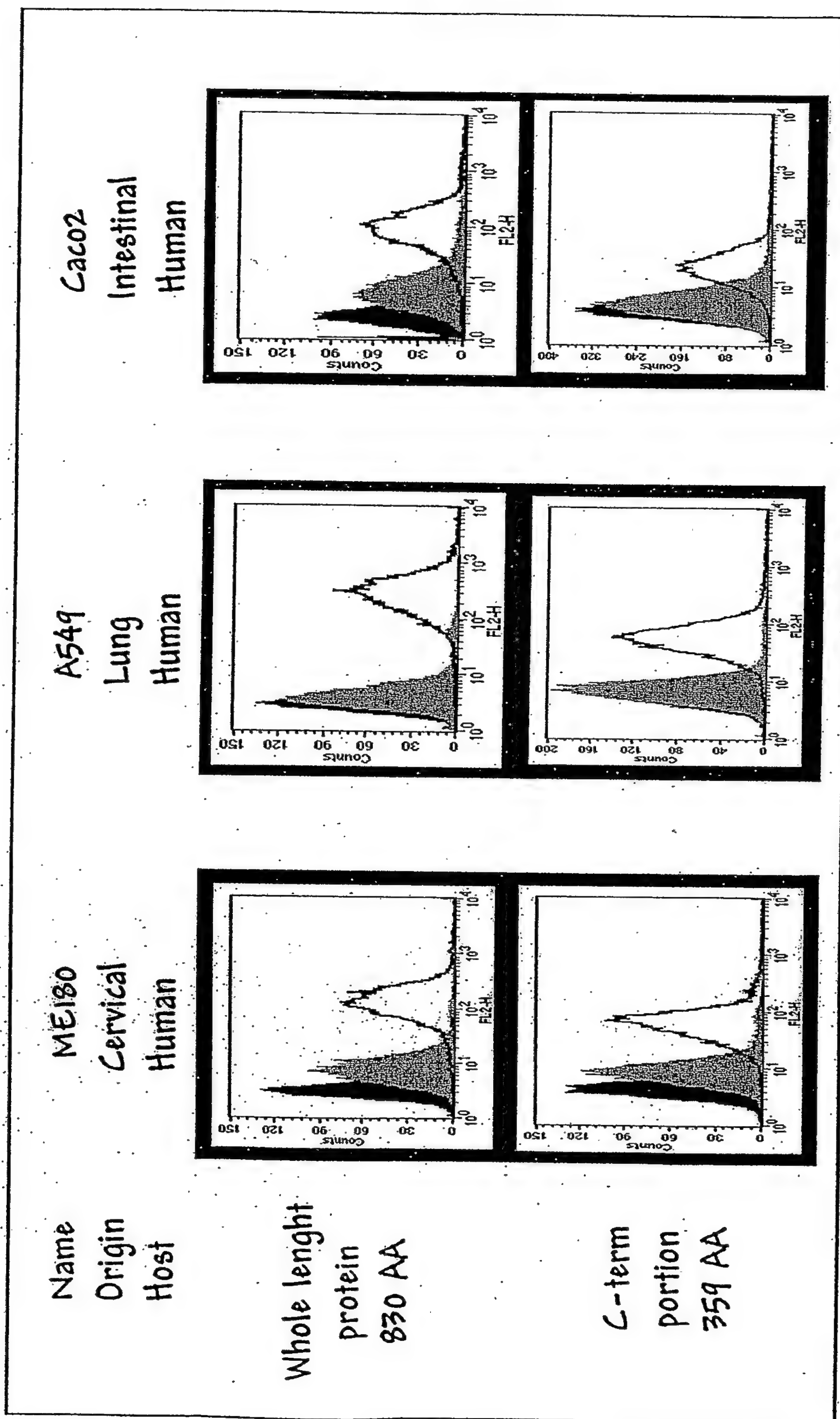
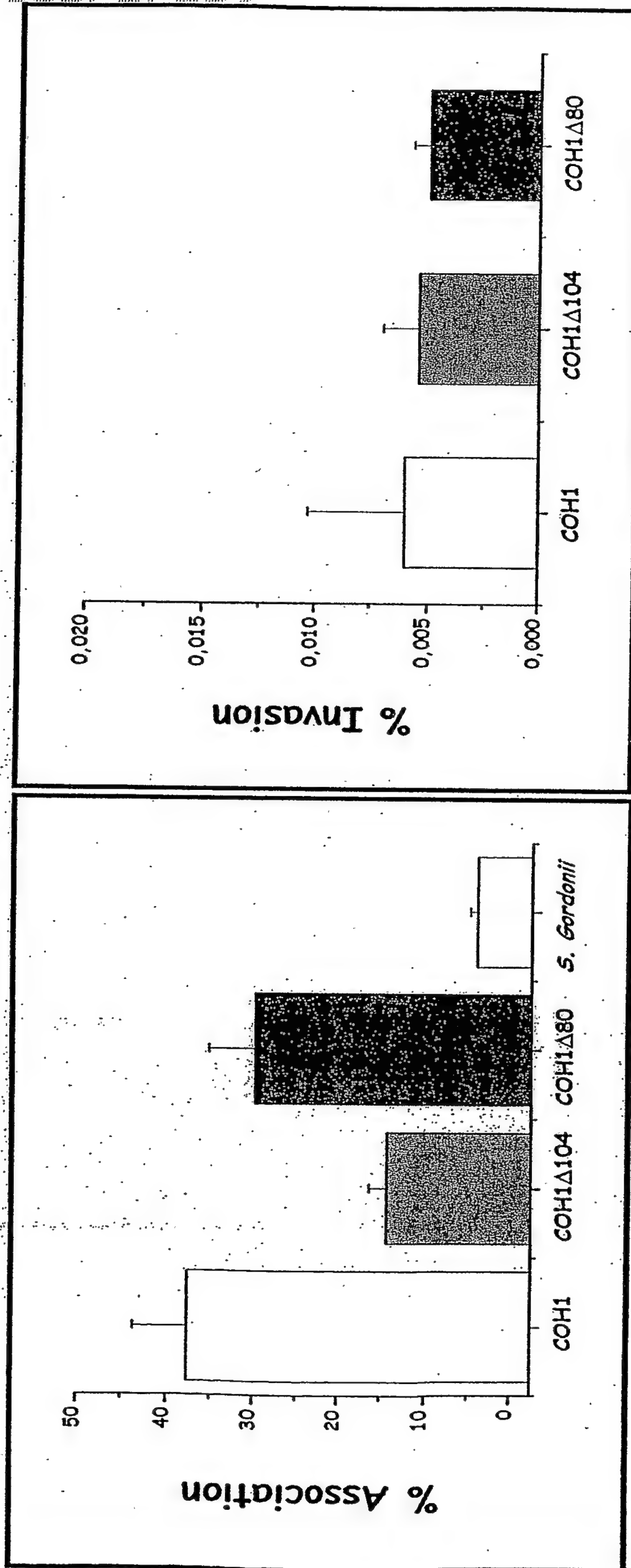


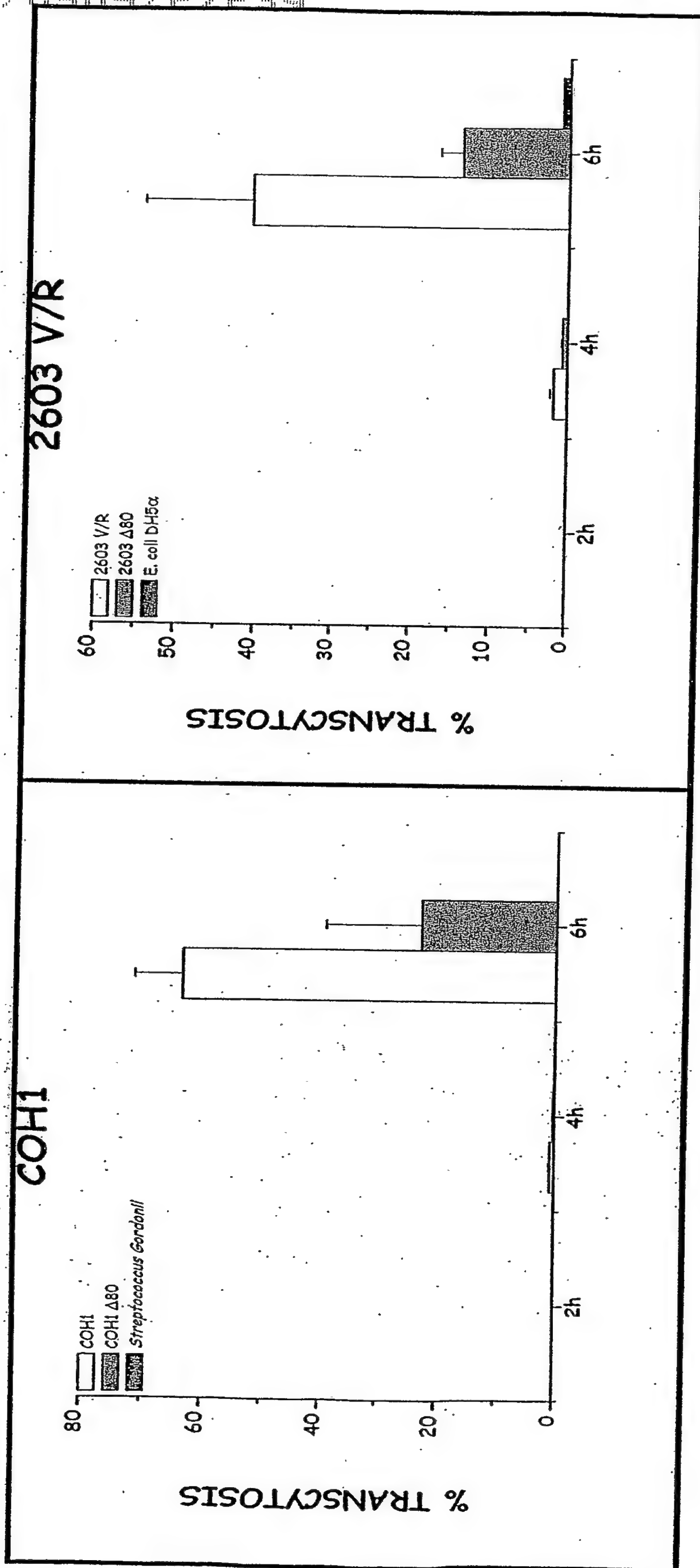


Figure 30



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Figure 31





PCT/US2005/027239/47/487

Figure 32

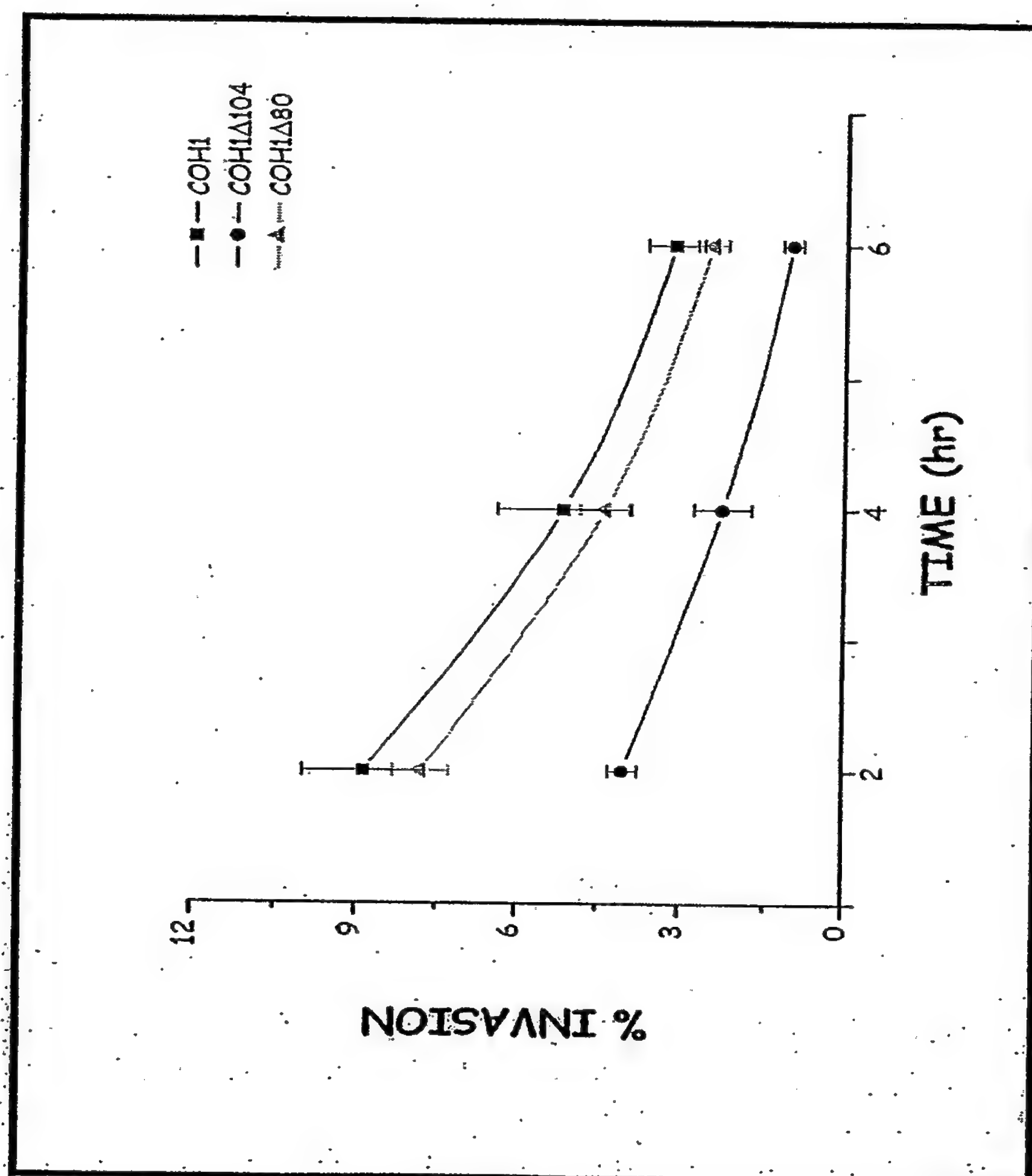
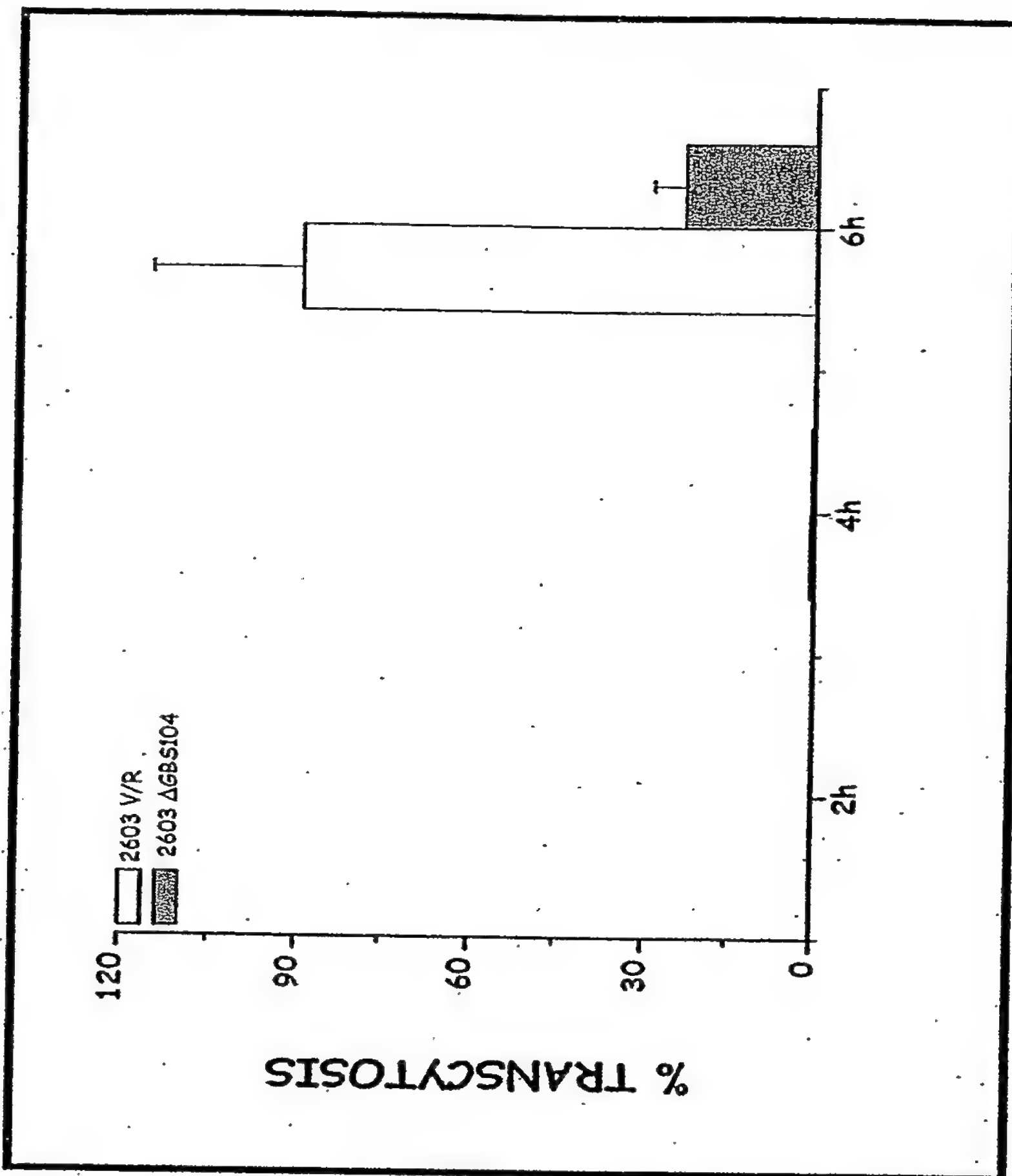
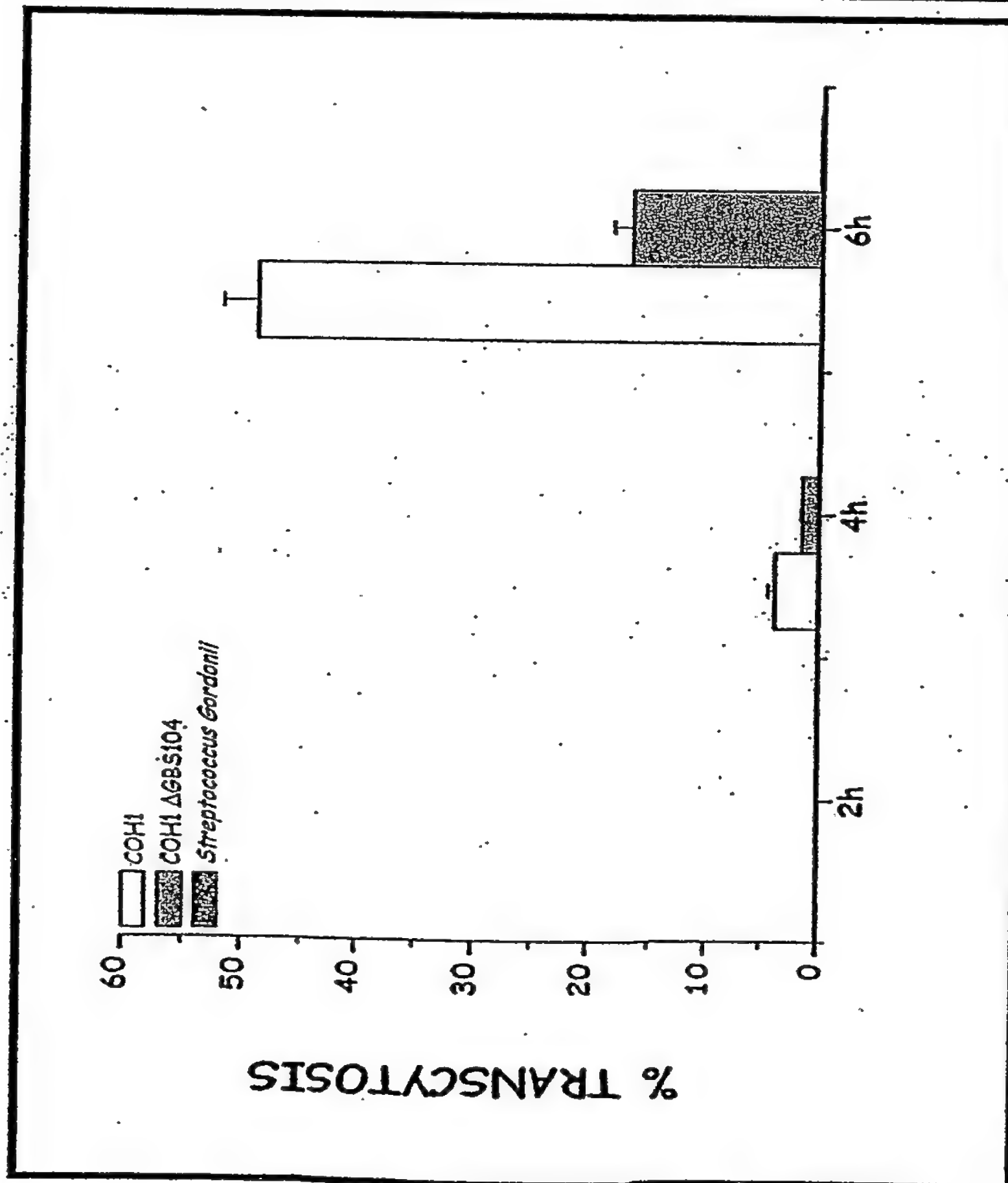


Figure 33

COH1

2603 V/R



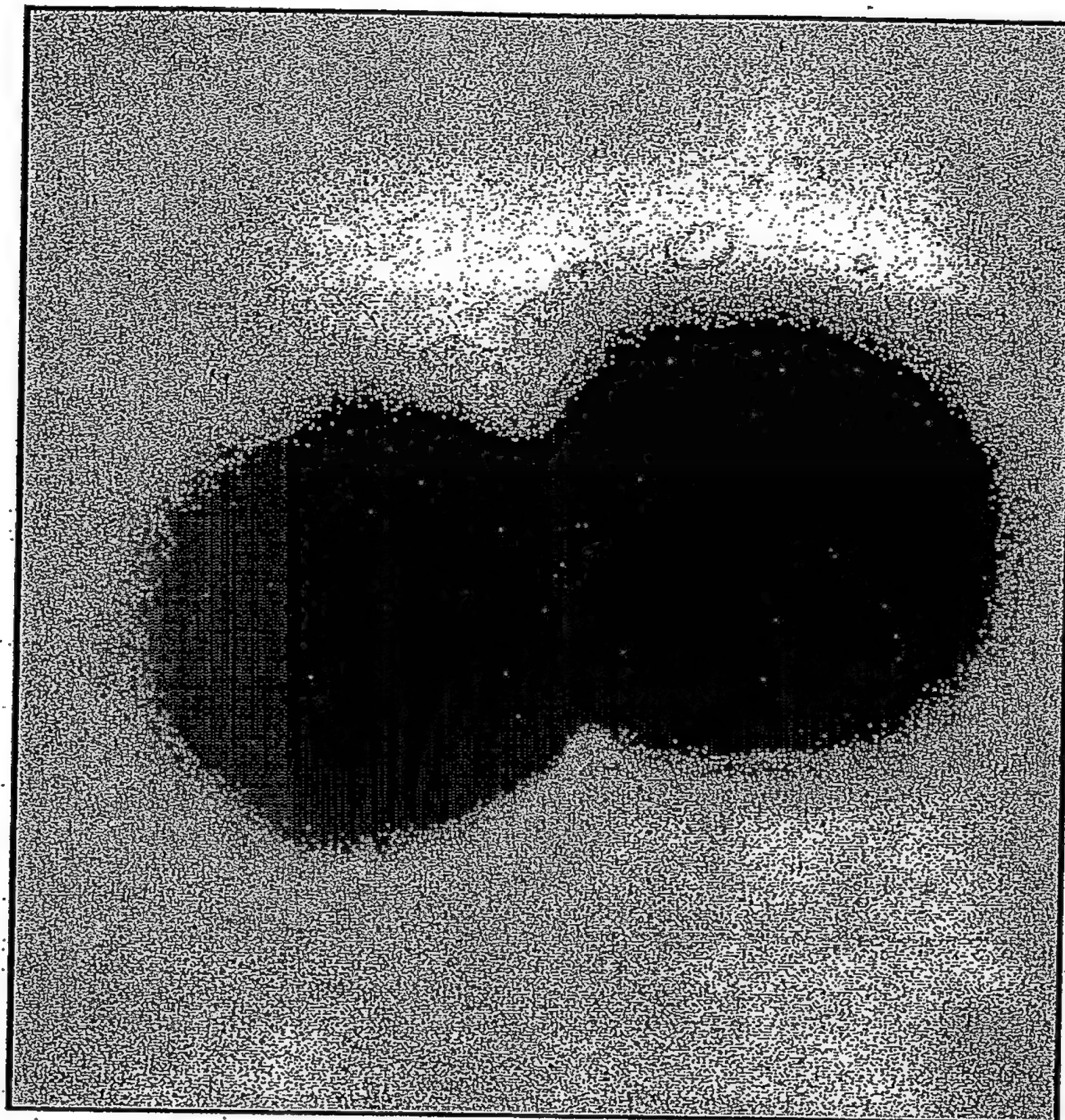
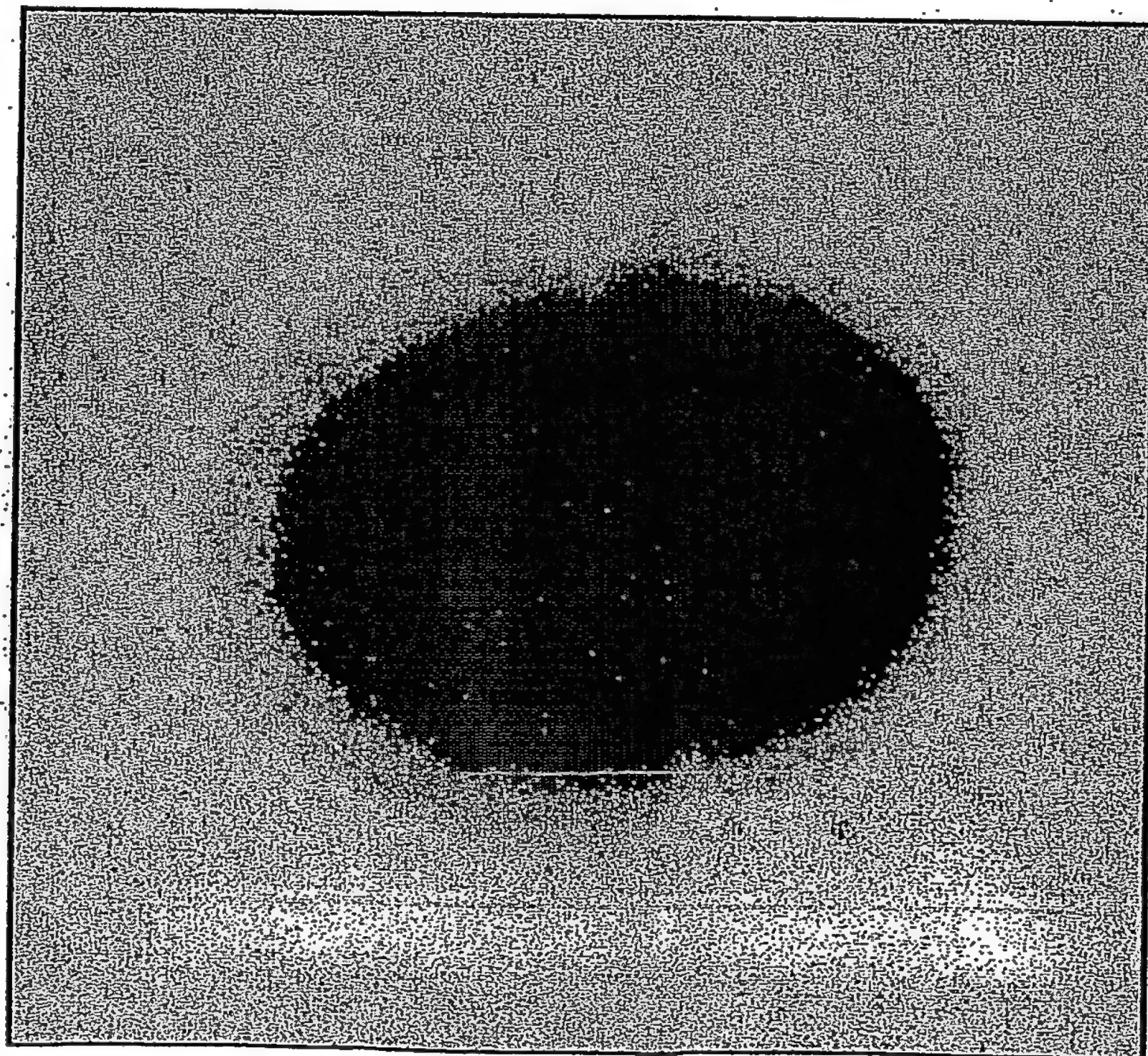


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# GBS STRAIN COH1 over GBS80

Figure 34

Negative staining EM





# GBS STRAIN COH1 over GBS80

IEM anti-GBS80 (gold particles 10nm)

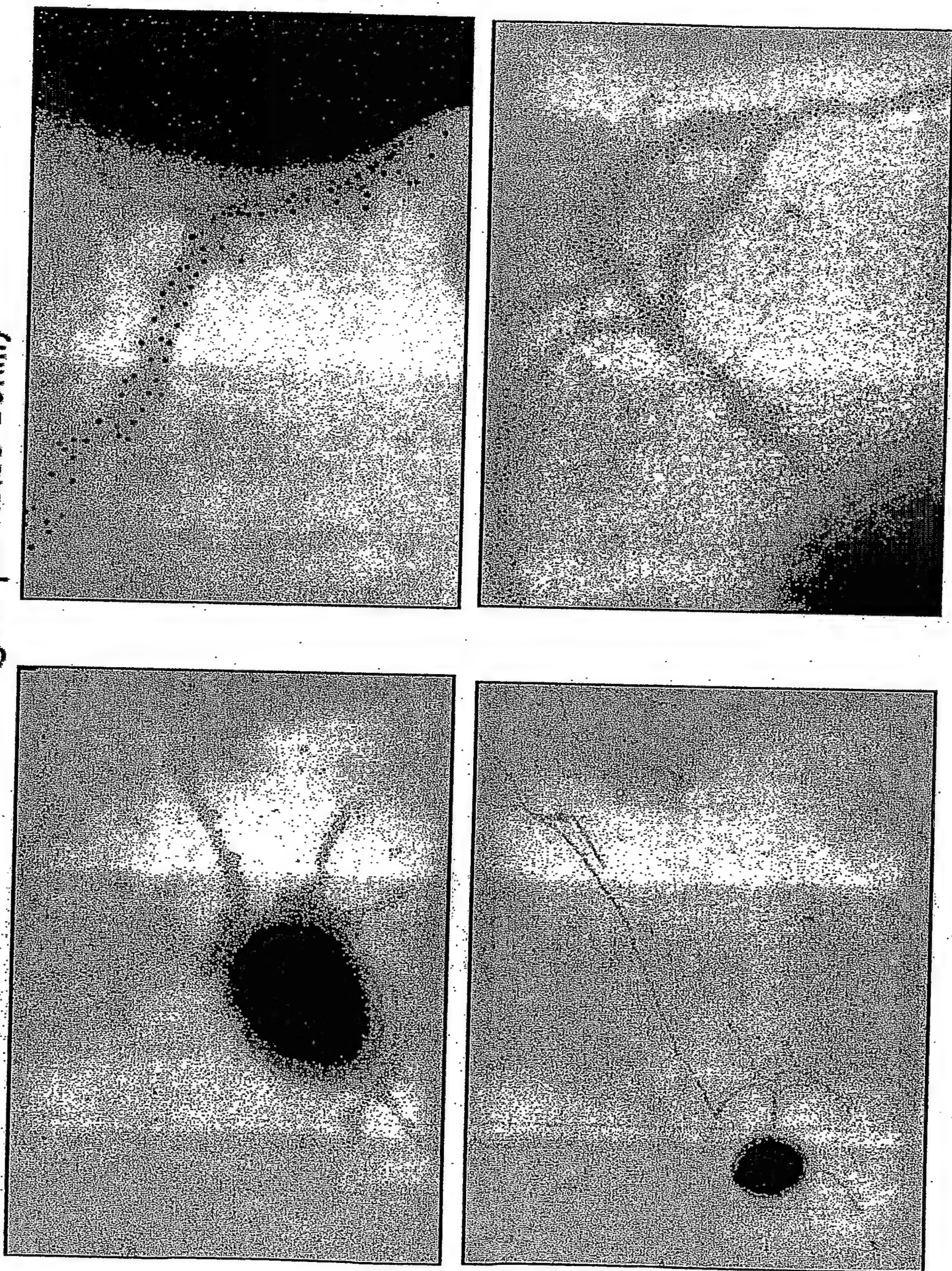


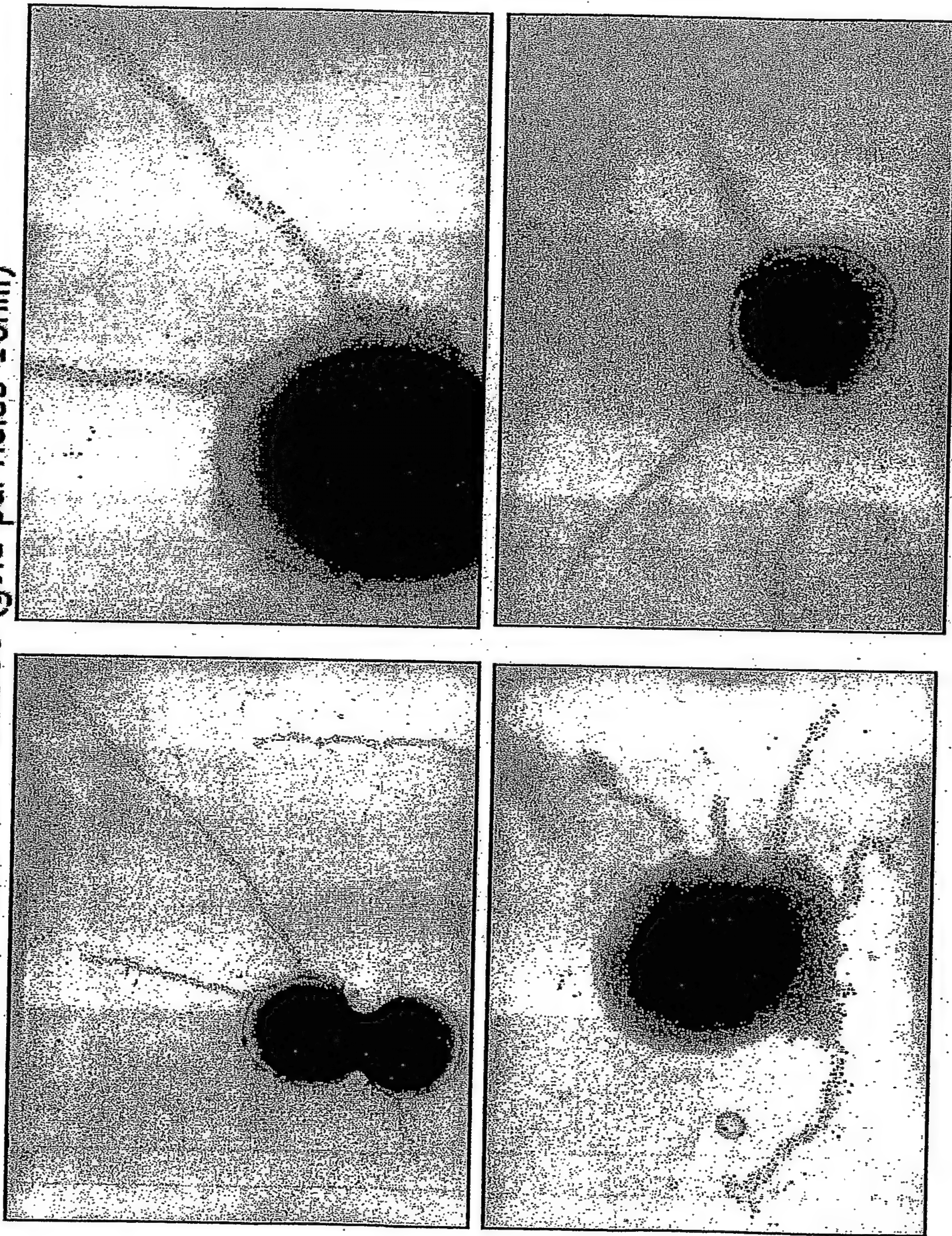
Figure 35



# GBS STRAIN COH1 over GBS80

Figure 36

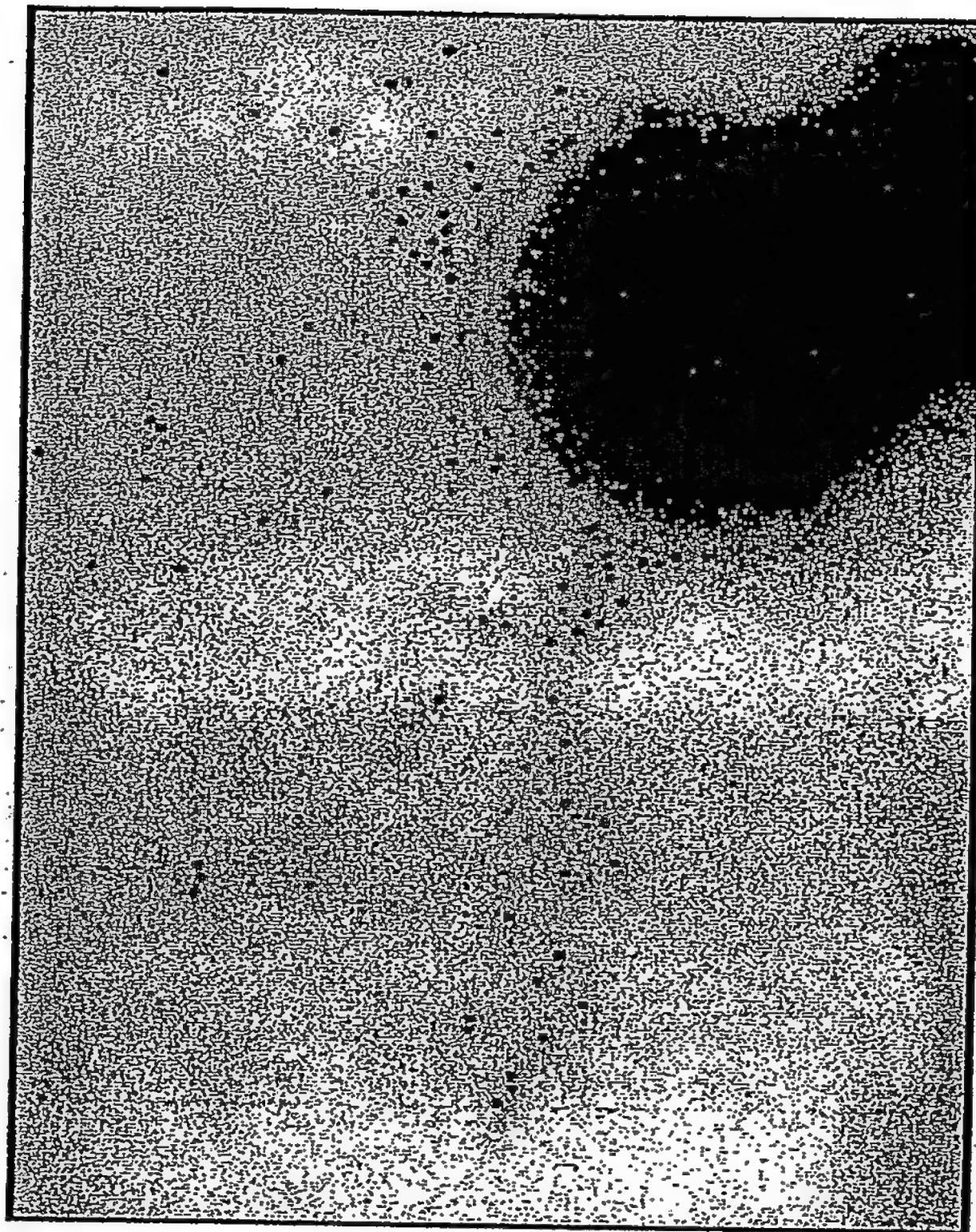
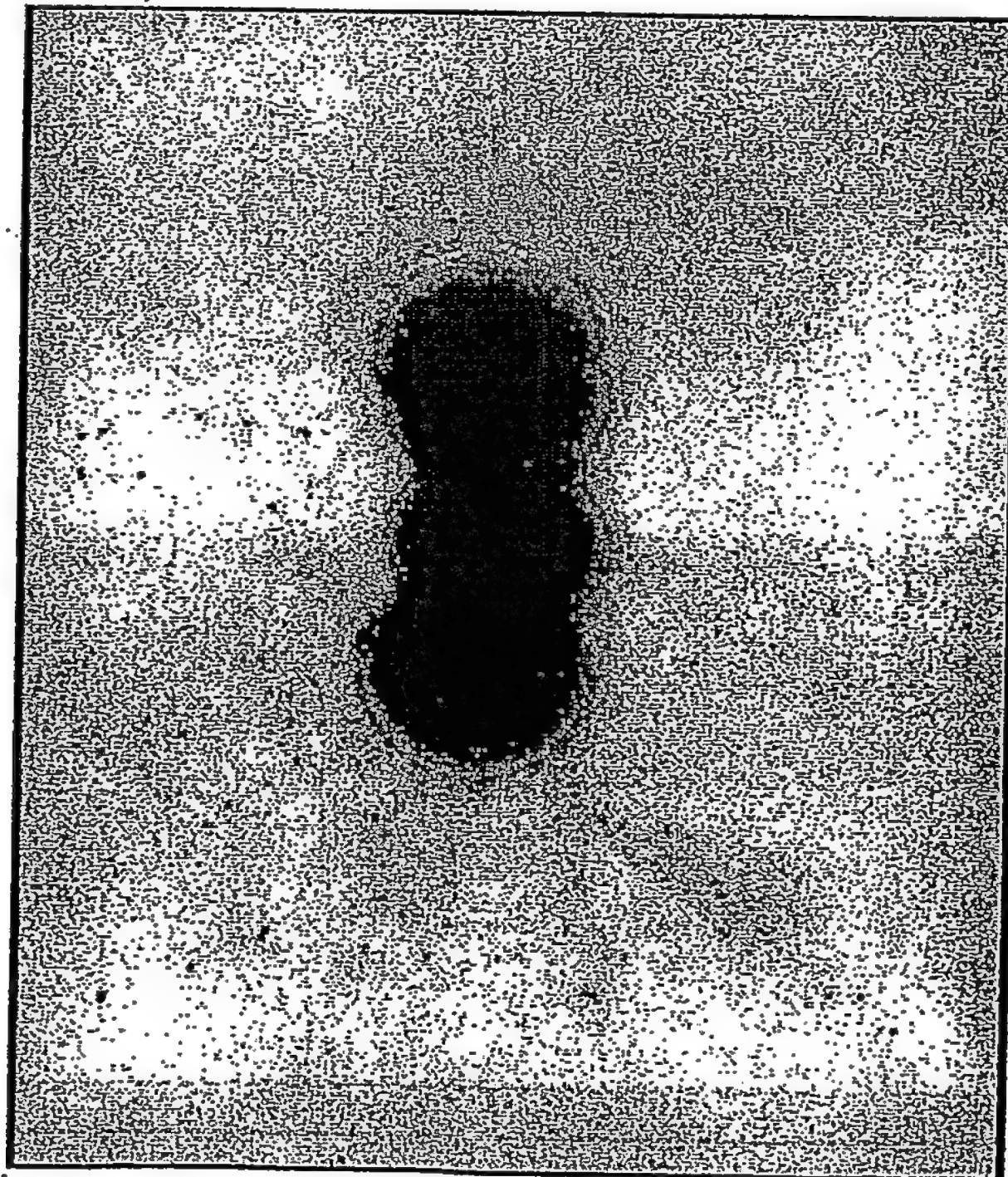
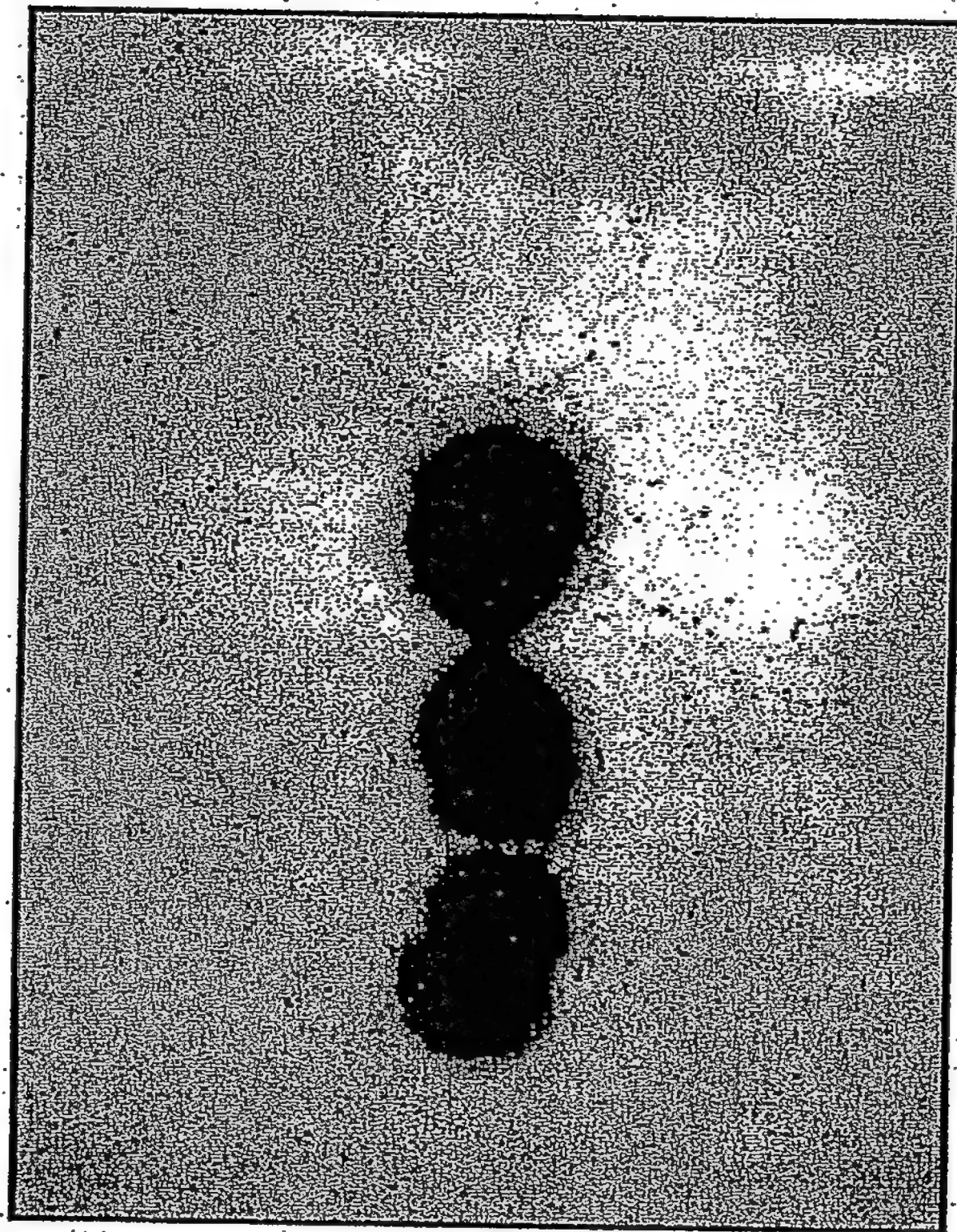
IEM anti-GBS80 (gold particles 10nm)





# GBS STRAIN COH1 over GBS80 IEM anti-GBS80 (gold particles 20nm)

Figure 37

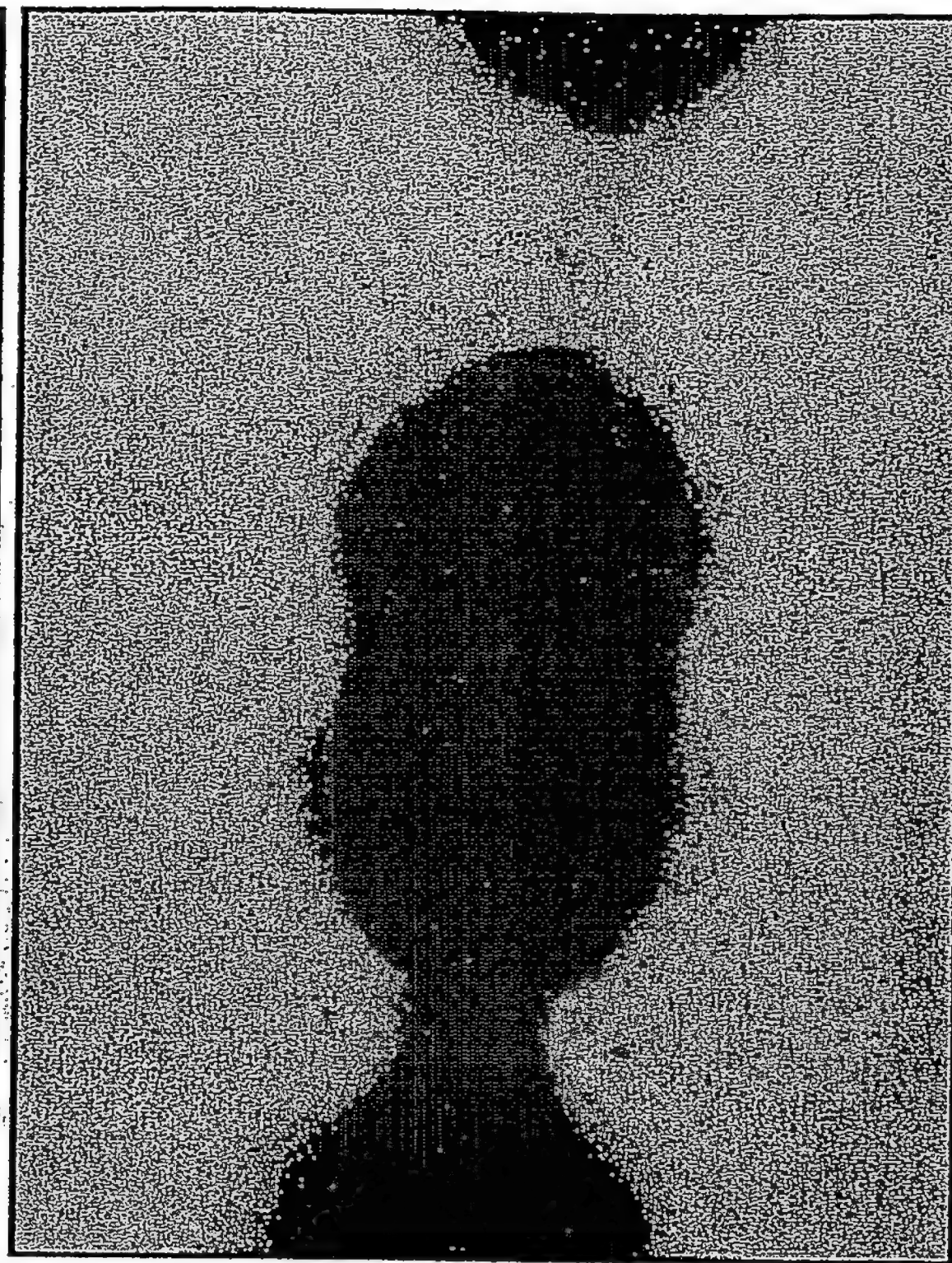
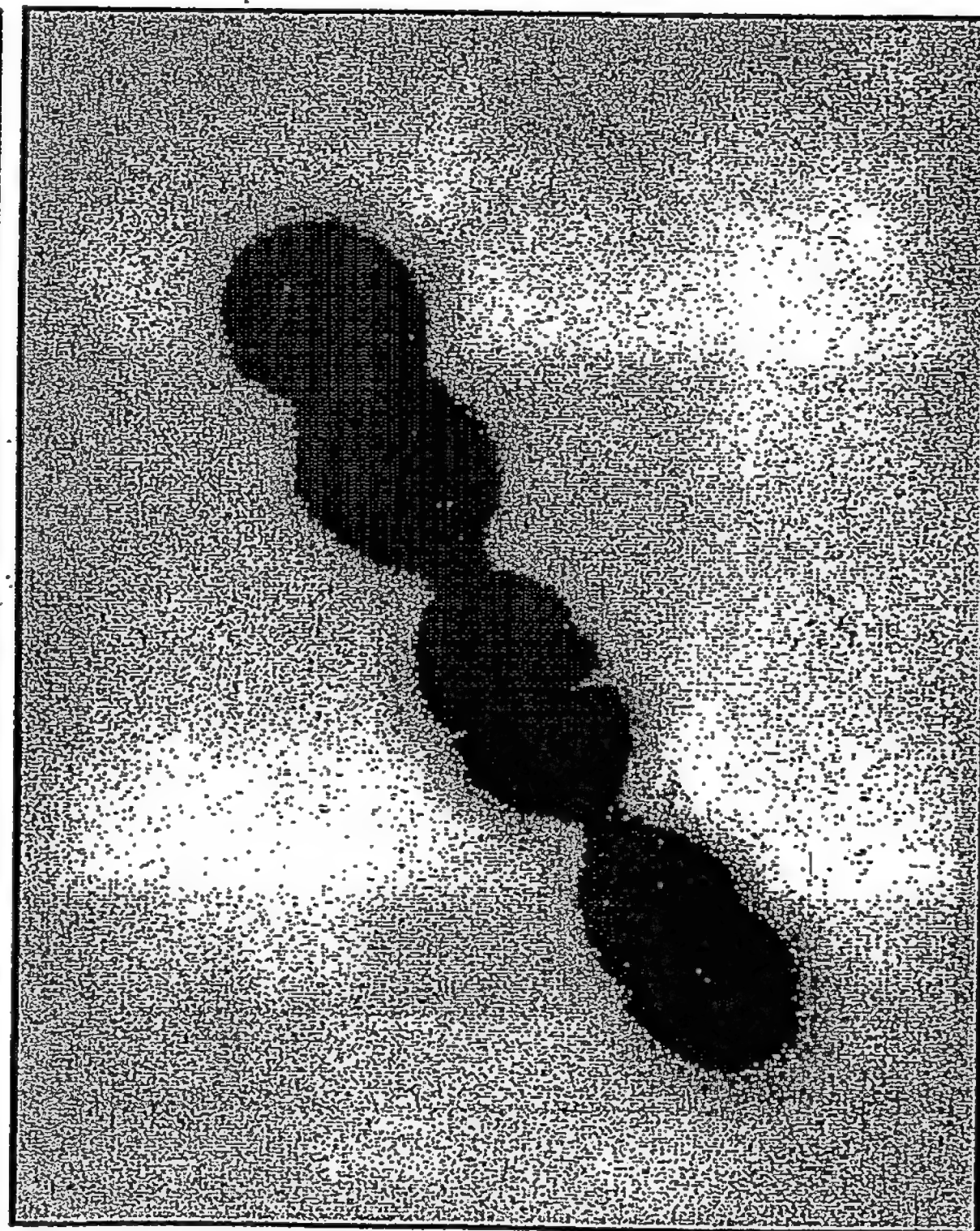
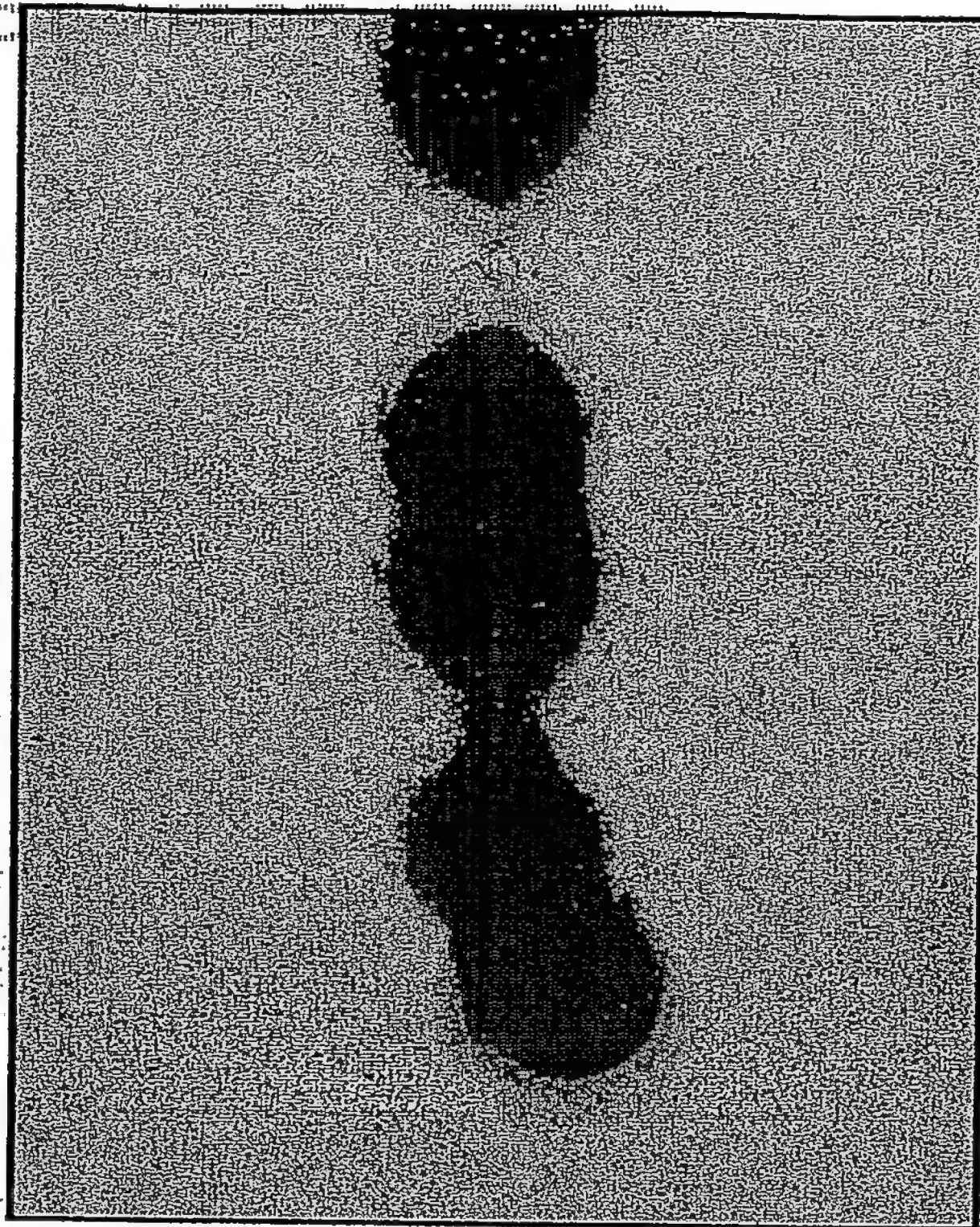
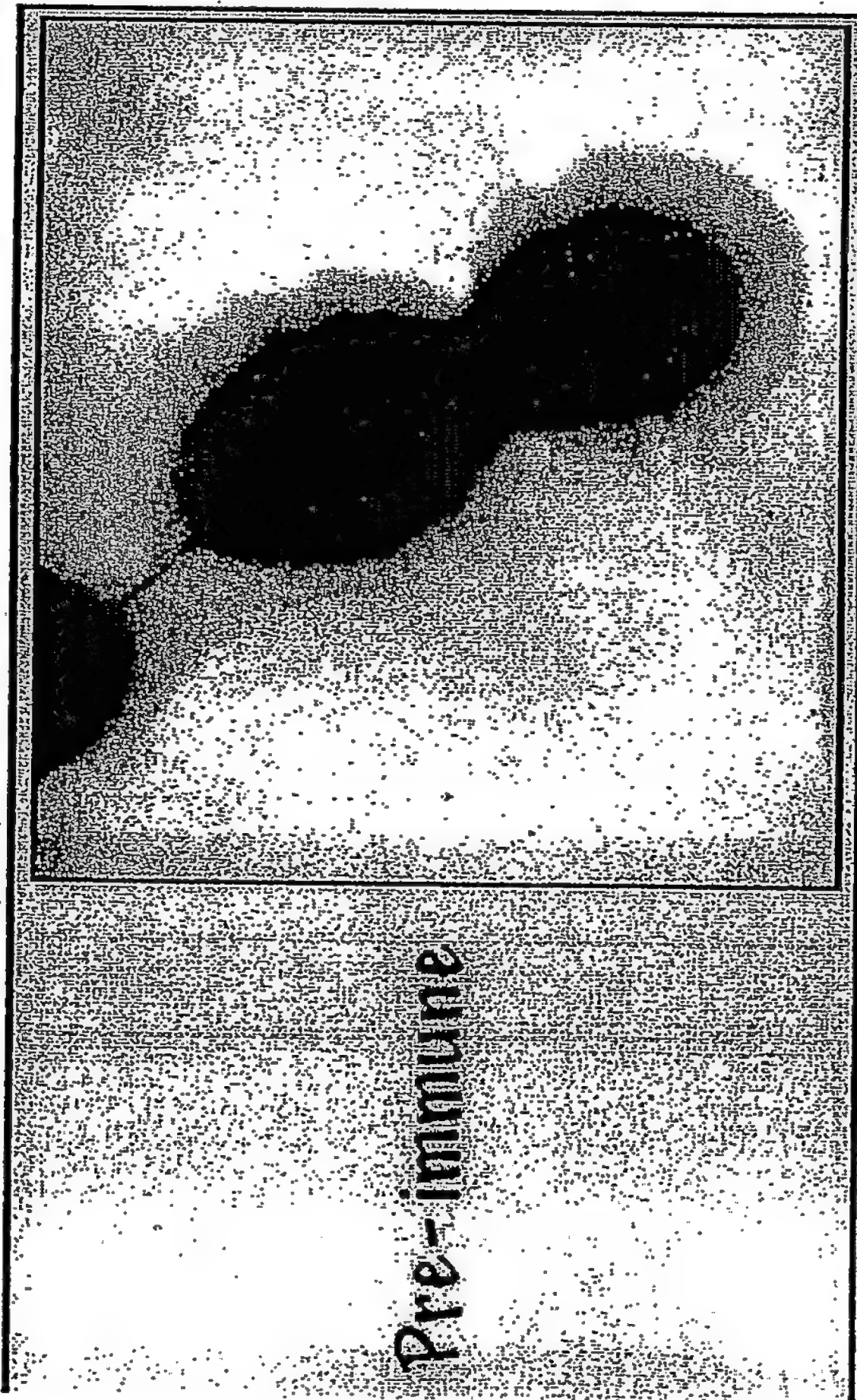




# GBS STRAIN COH1 over GBS80

Figure 38

IEM anti-GBS104 (gold particles 10nm)





# GBS STRAIN COH1 over GBS80

Figure 39

IEM anti-GBS80 (gold particles 20nm) anti-GBS104 (gold particles 10nm)

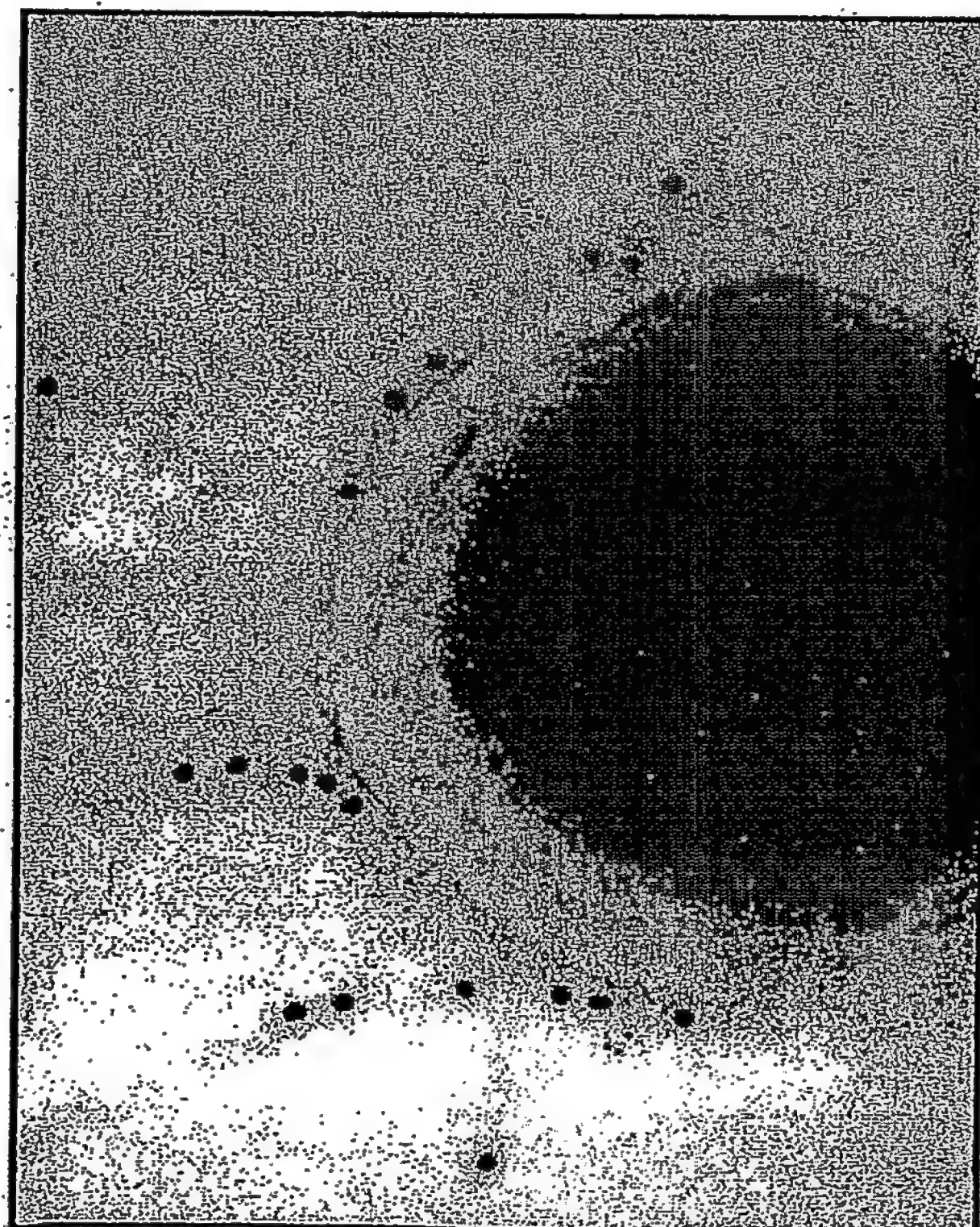
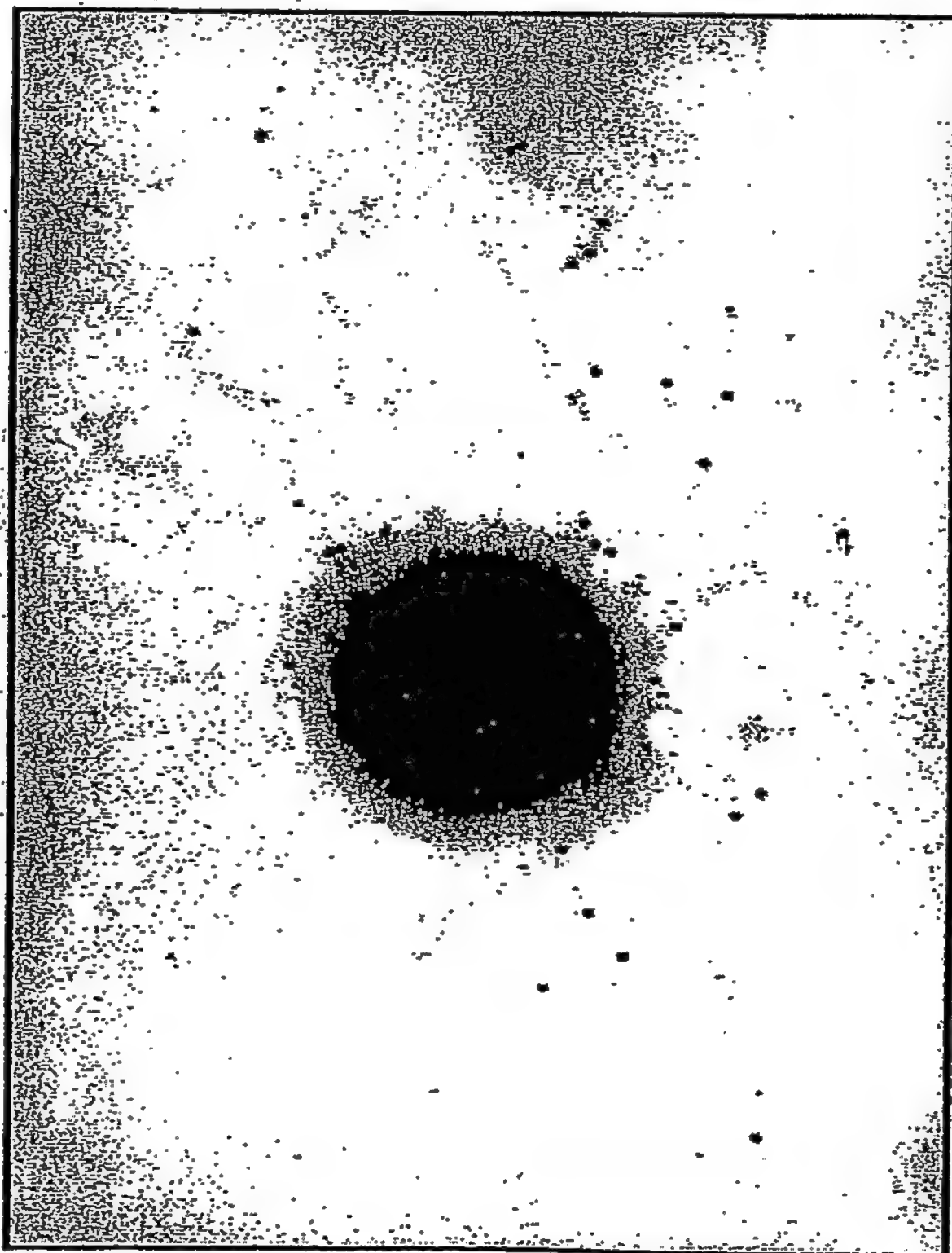
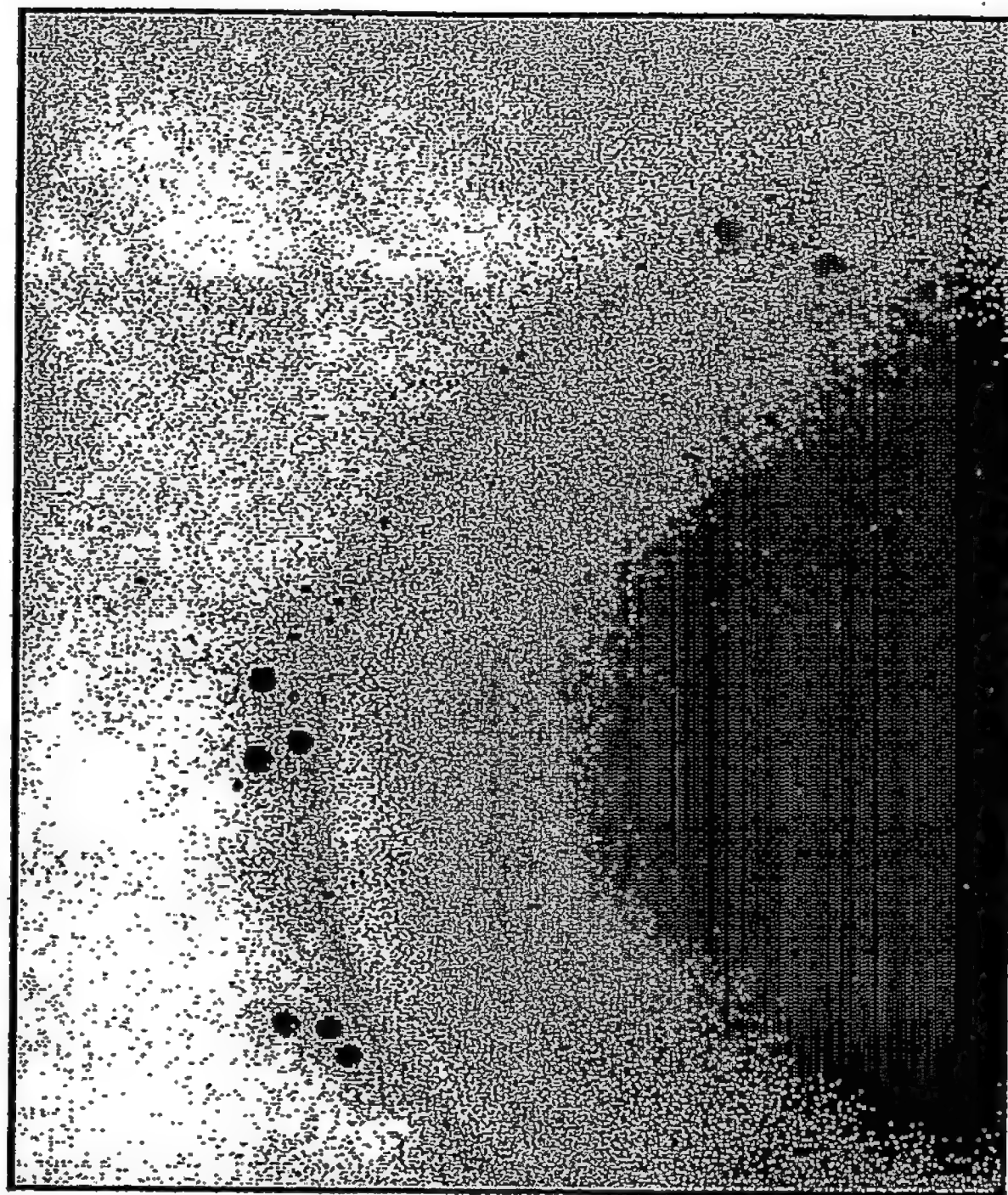
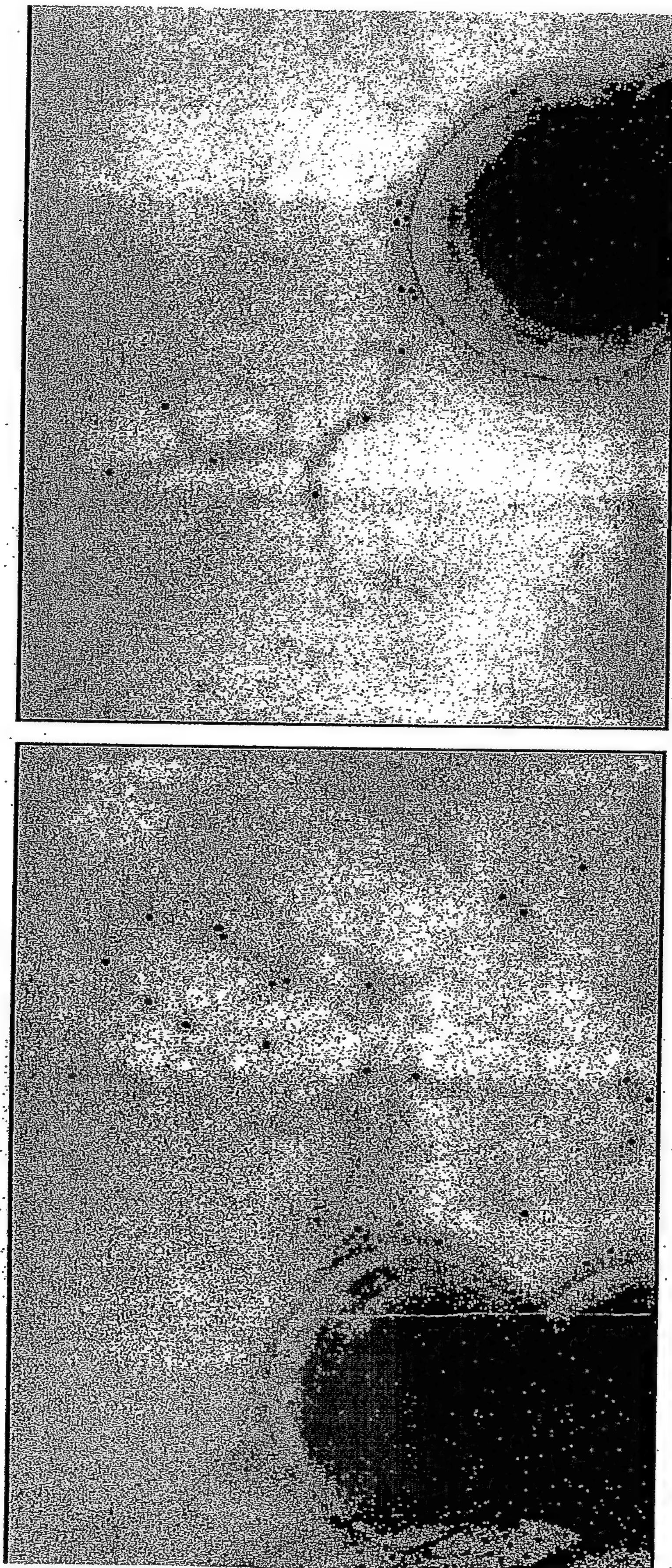




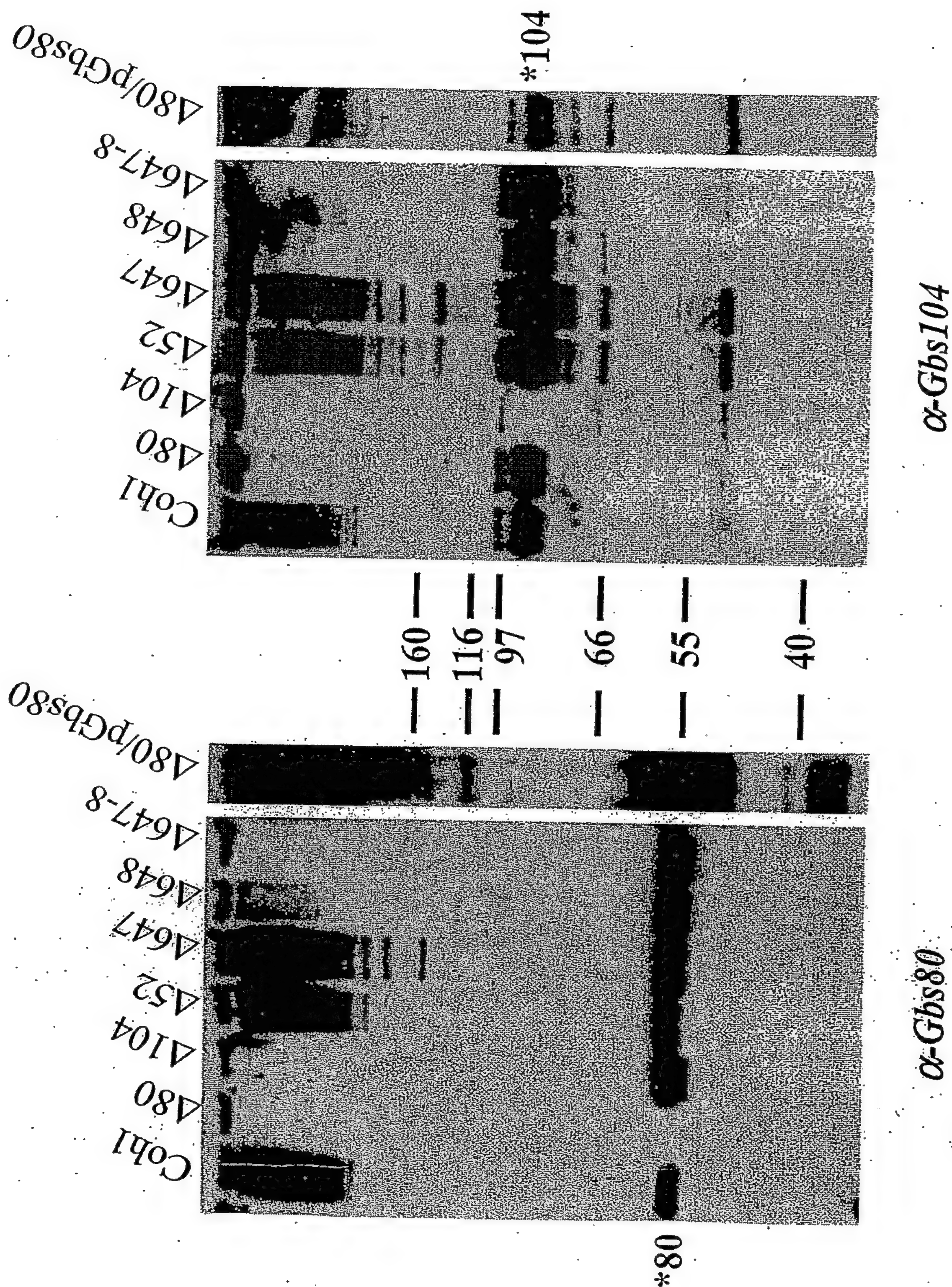
Figure 40

**GBS STRAIN COH1 over GBS80****IEM anti-GBS80 (gold particles 20nm) anti-GBS104 (gold particles 10nm)**



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**Figure 41: GBS 80 is necessary for polymer formation, GBS104 and sortase SAG0648 are necessary for efficient assembly**



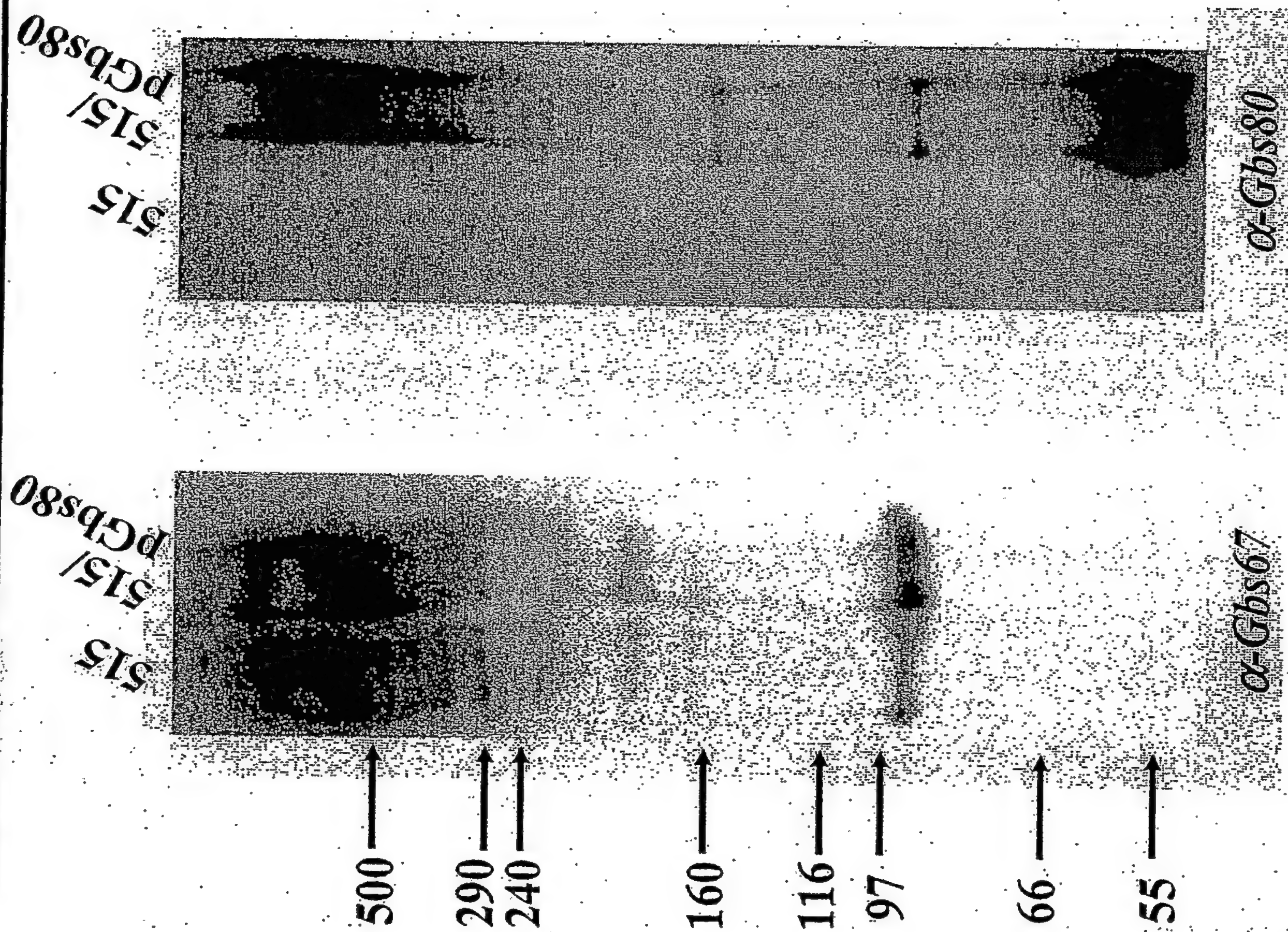


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Figure 42: Gbs67 is part of a second pilus;

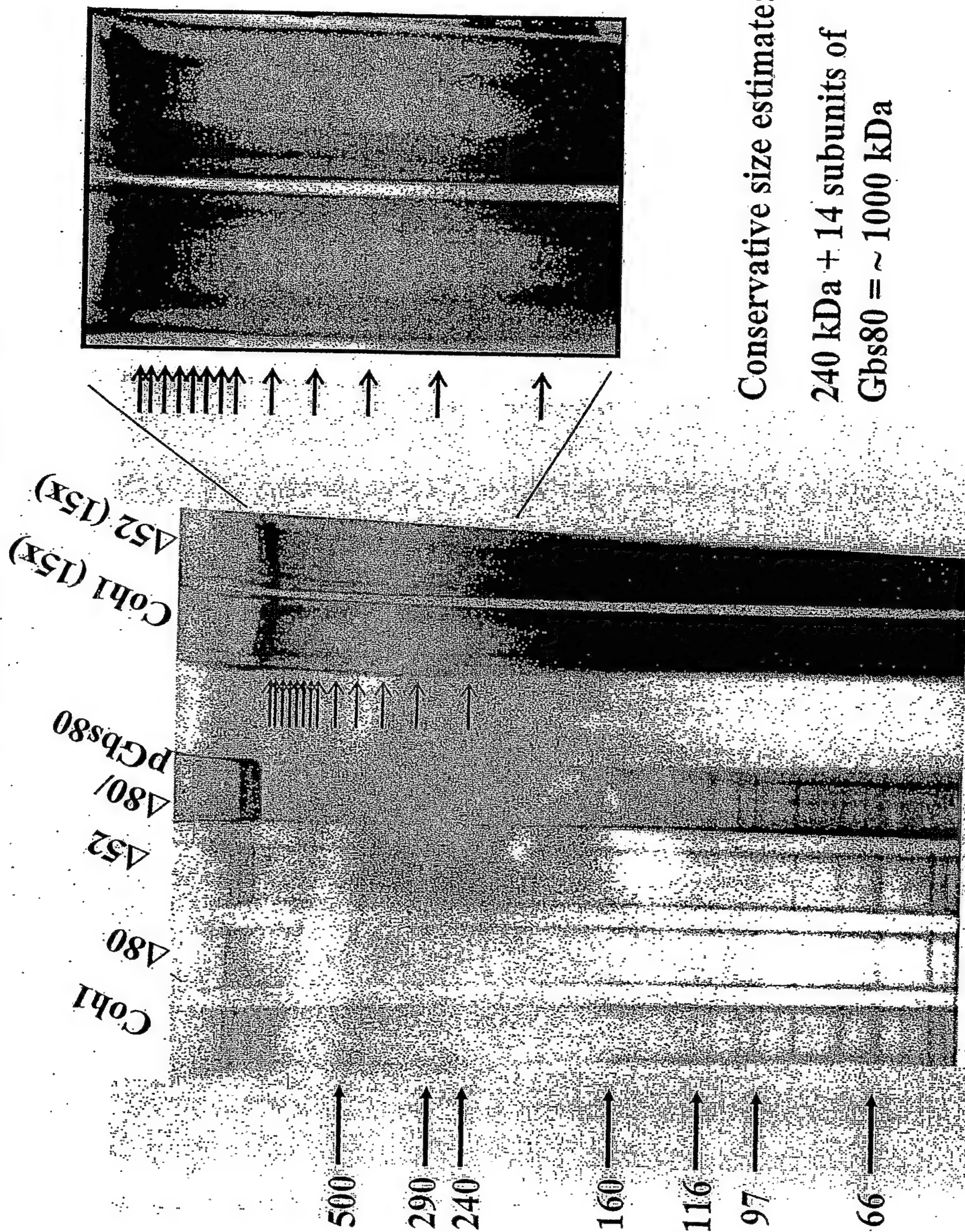
Gbs80 is polymerized in strain 515

(515 lacks sortase 647-8, but has AI-2 sortases)





**Figure 43: Two macro-molecules are visible in Coh1 at >1000 kDa, one is the Gbs80 pilin**





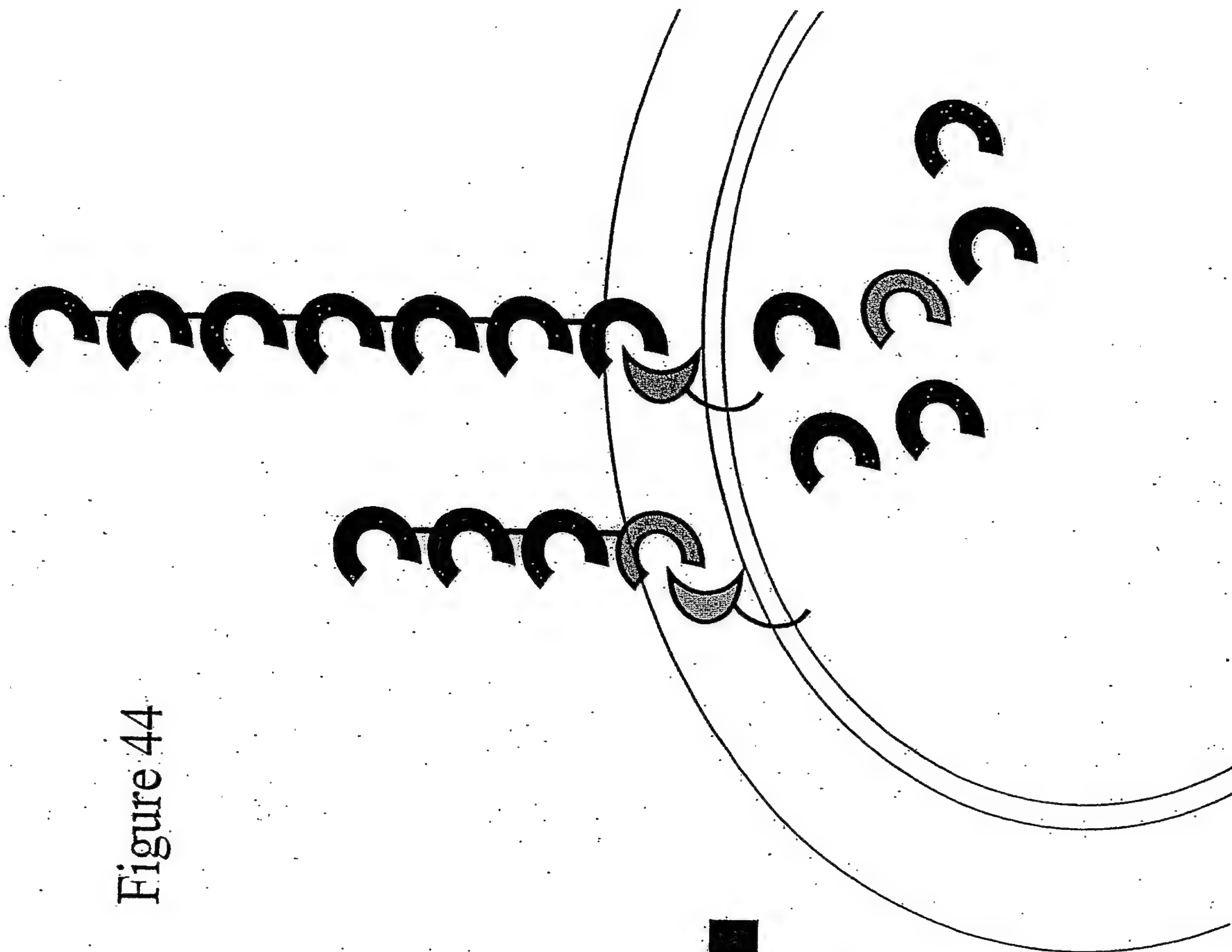


Figure 44

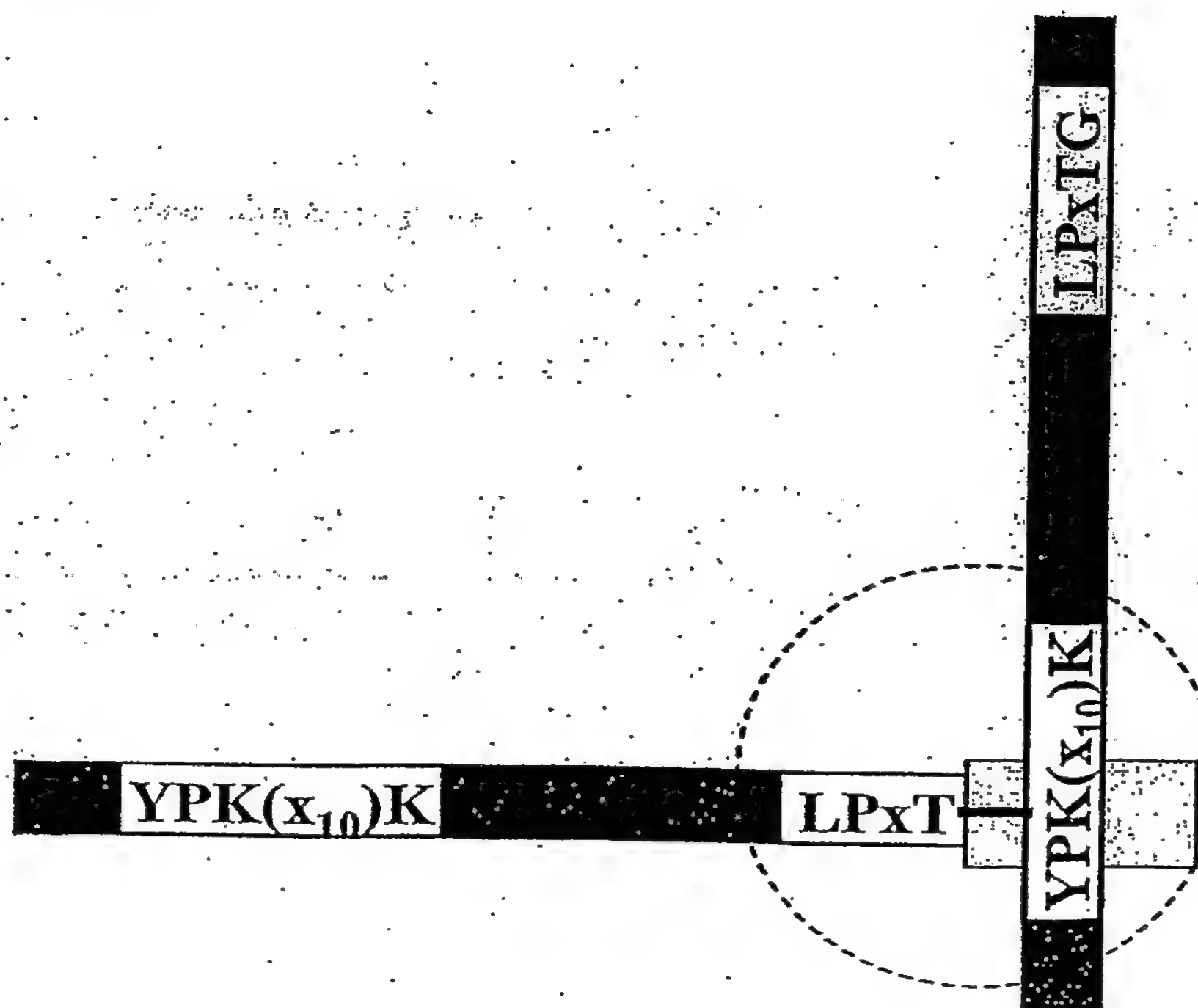
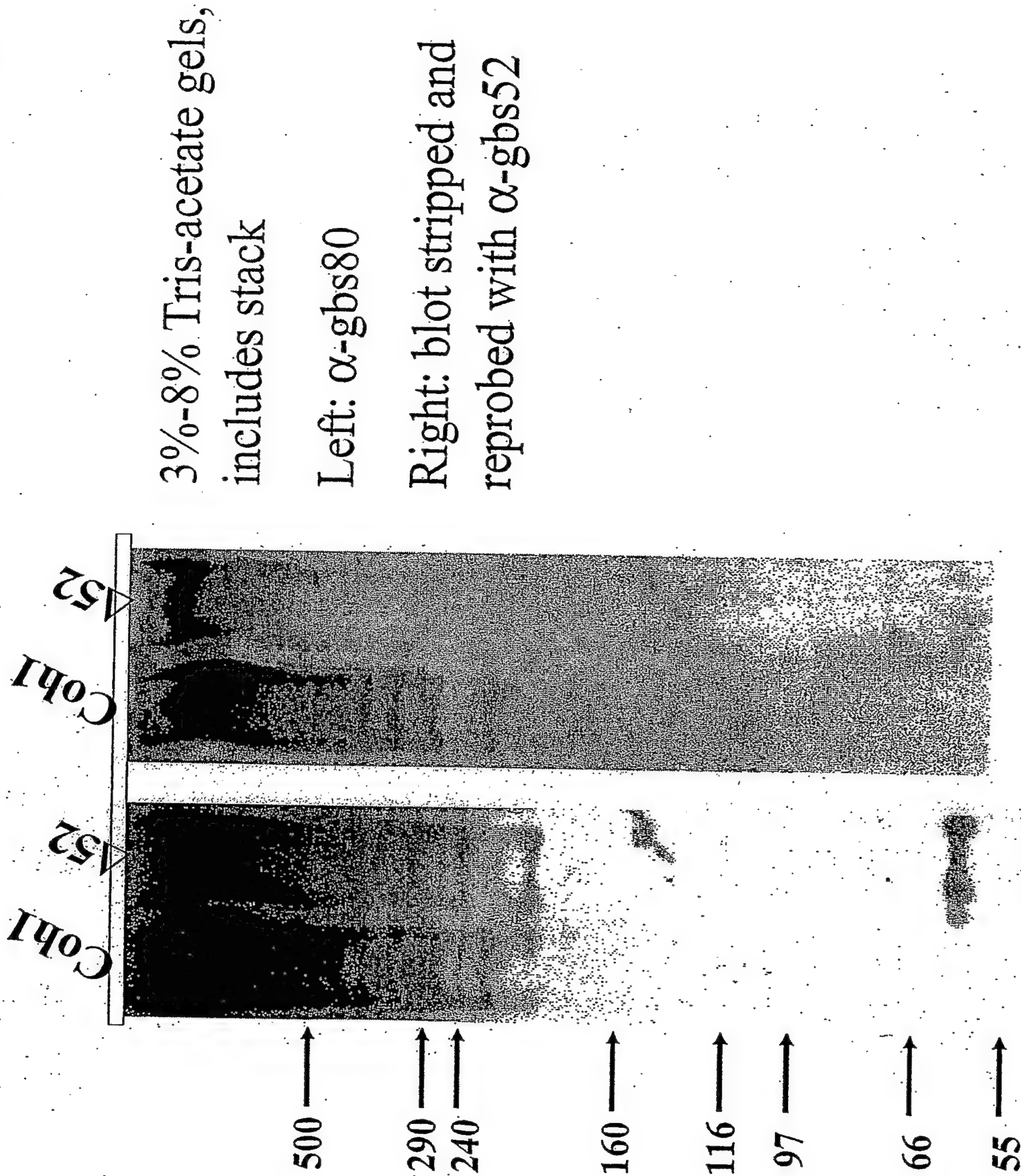
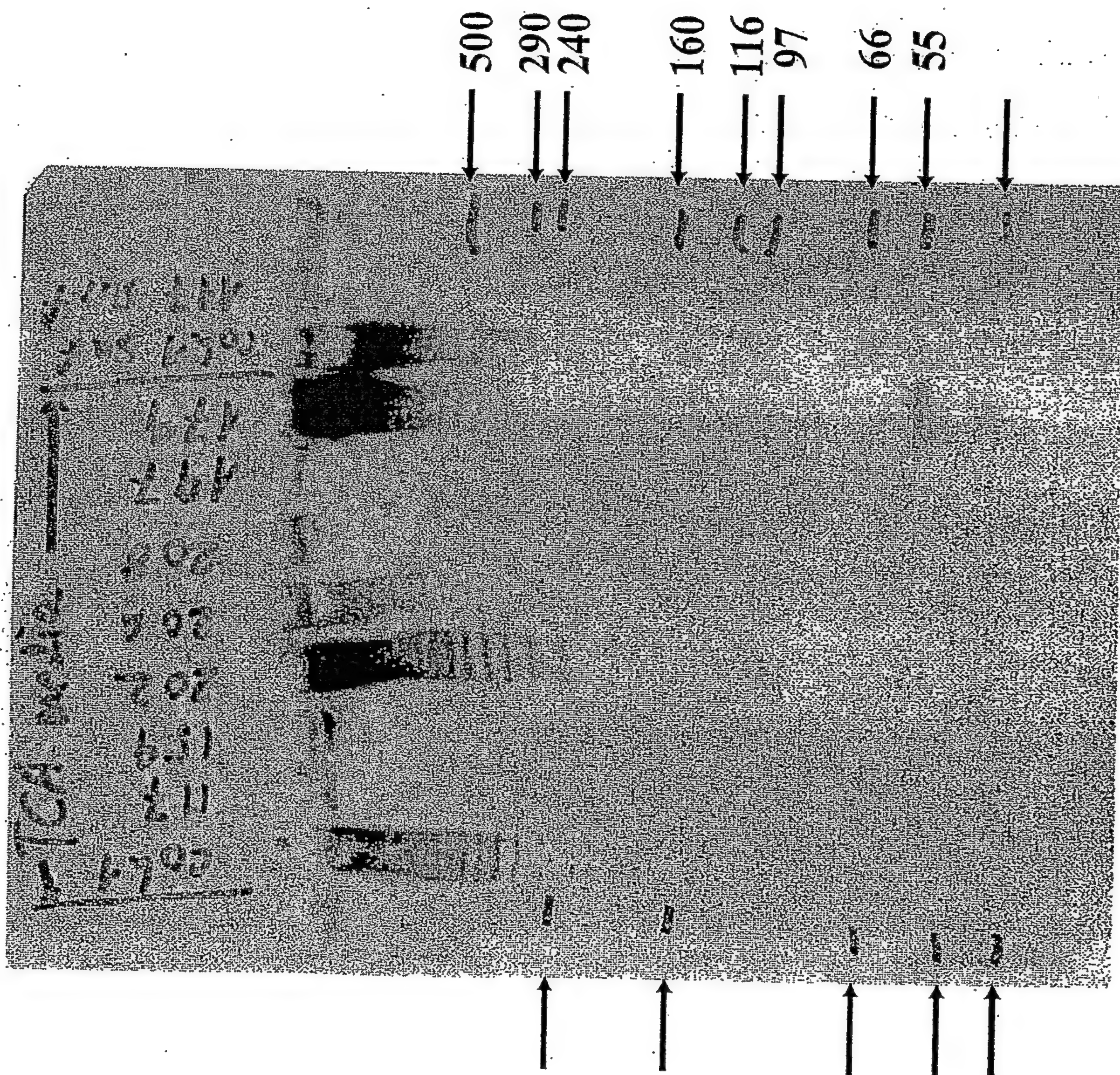


Figure 45: Gbs52 is a minor component of the GBS pilus



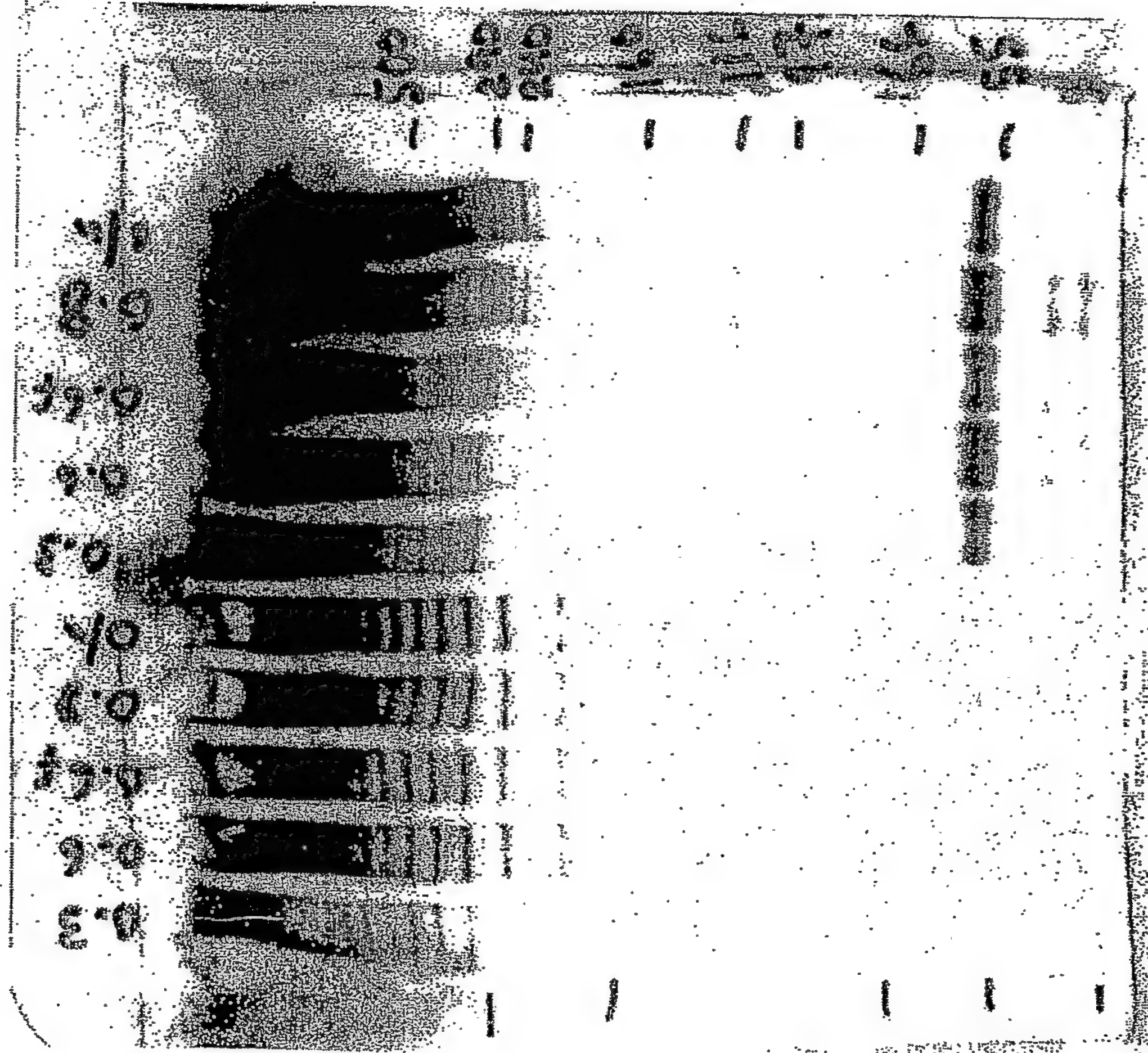


**Figure 46: The pilus is found in the supernatant of the bacterial culture**





**Figure 47: The pilus is found in the supernatant  
of cultures in all growth phases**



TCA precipitation of 1 ml  
of THB culture supernatant  
run on 3-8% SDS-PAGE.  
OD600 nm are noted above  
samples, "f" indicates  
supernatant was filtered  
(0.2  $\mu$ M syringe filter).

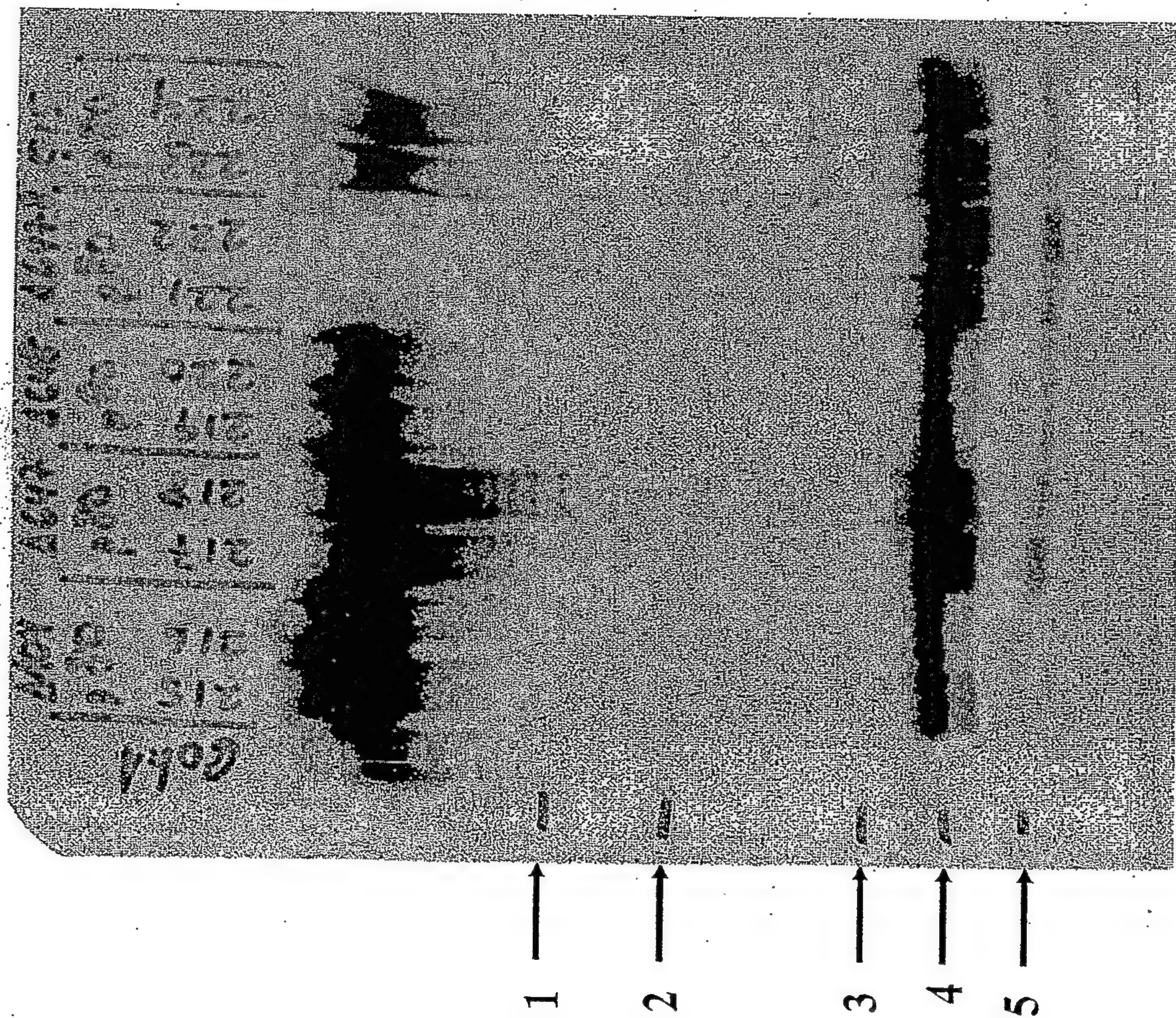
Left five samples: Coh1.

Right five samples: 179  
( $\Delta$ Gbs80/pGbs80).



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**Figure 48: In Coh1, only the gbs80 protein and one sortase (sag0647 or sag0648) is required for polymerization**



Over expression of gbs80 in various strain backgrounds (two clones each).

Total protein extract preparations.

Only the double sortase mutant does not polymerize gbs80.

Gbs80 is polymerized in the DK515 strain background (lacks adhesin island 1, adhesin island 2 is 2603-like). Presumably, sag1405&sag1406 are responsible for polymerization.



GBS STRAIN JM9030013  
IEM anti-GBS80

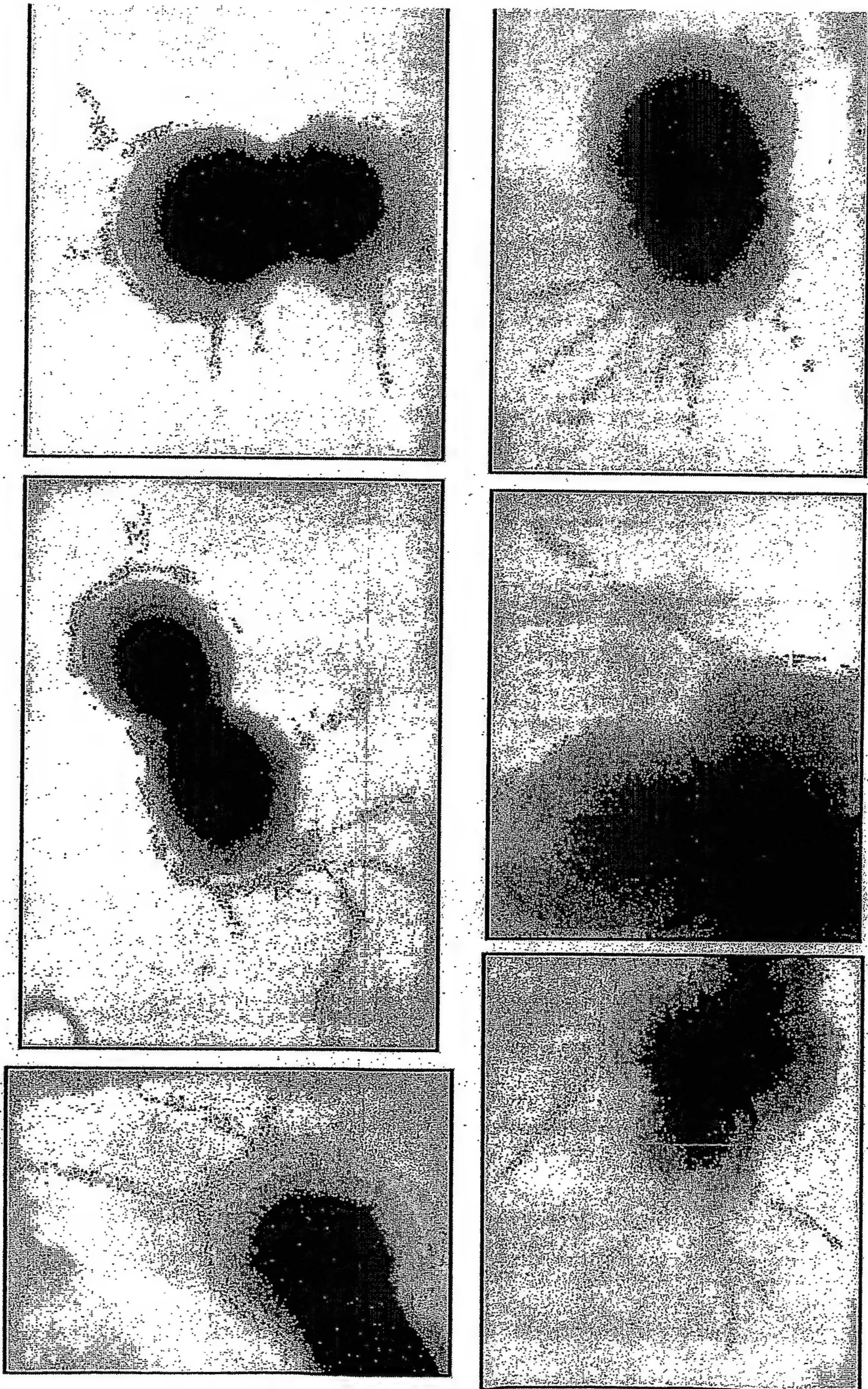


FIGURE 49



GBS STRAIN JM9030013

IEM anti-GBS104

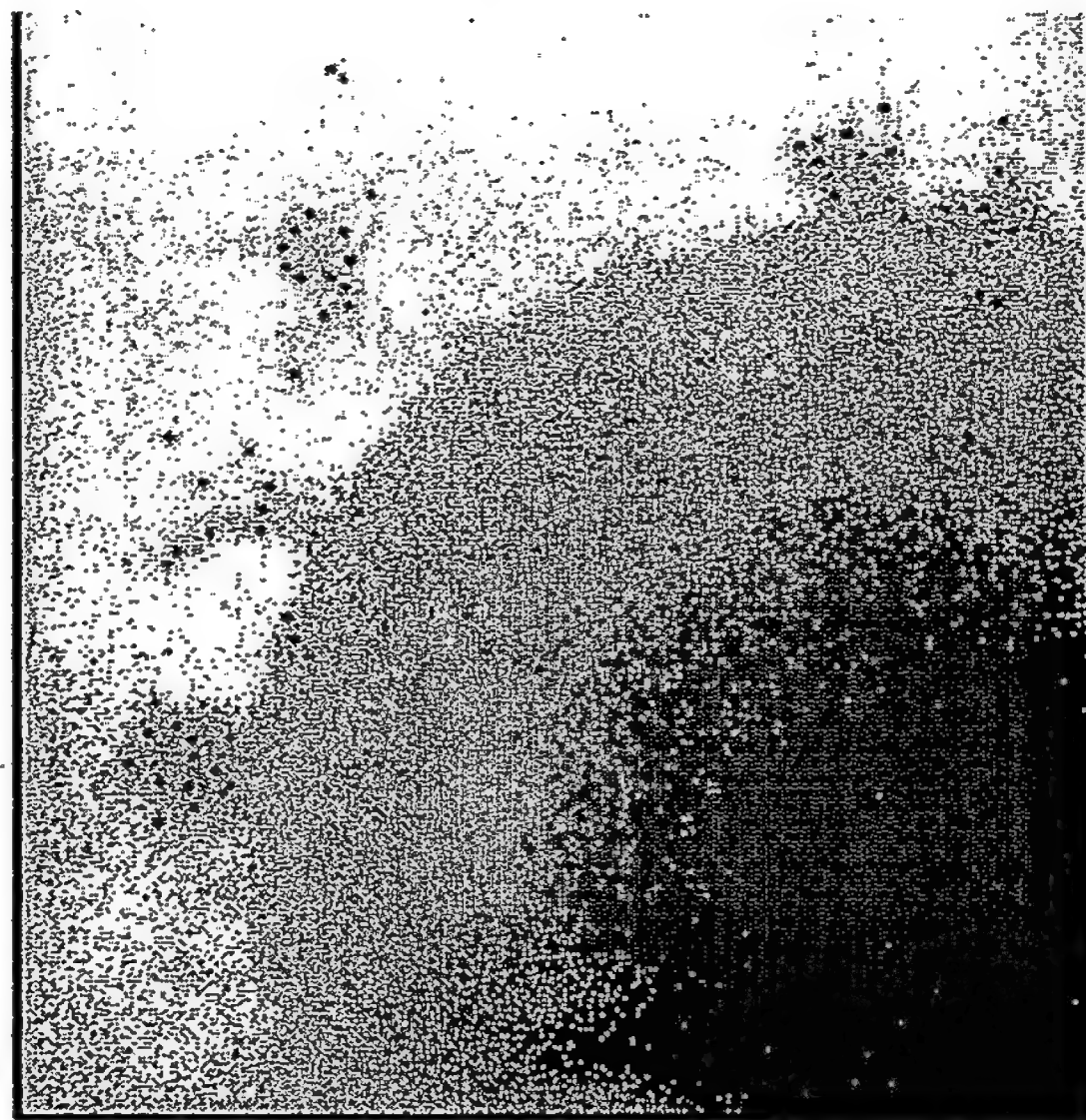
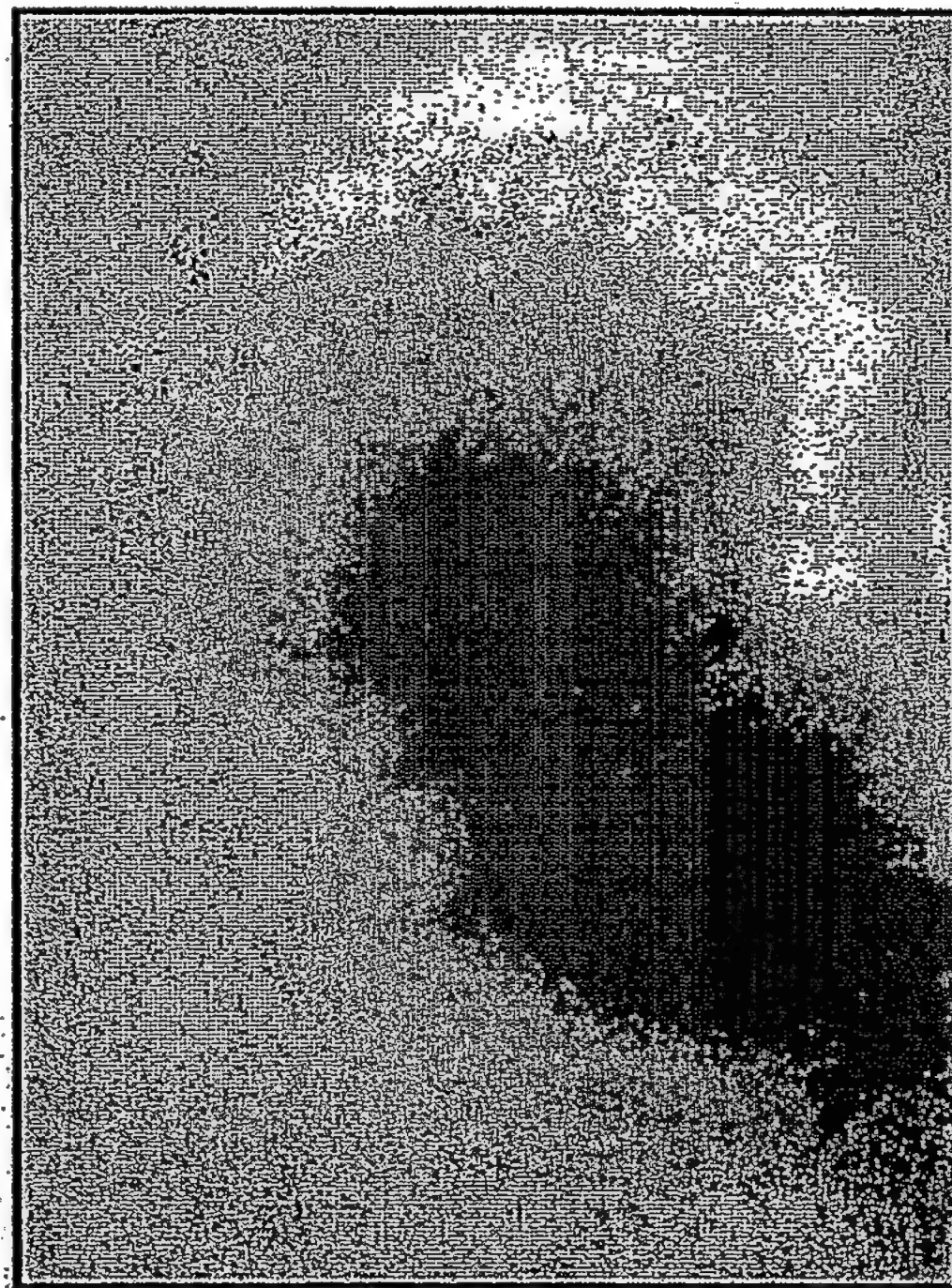
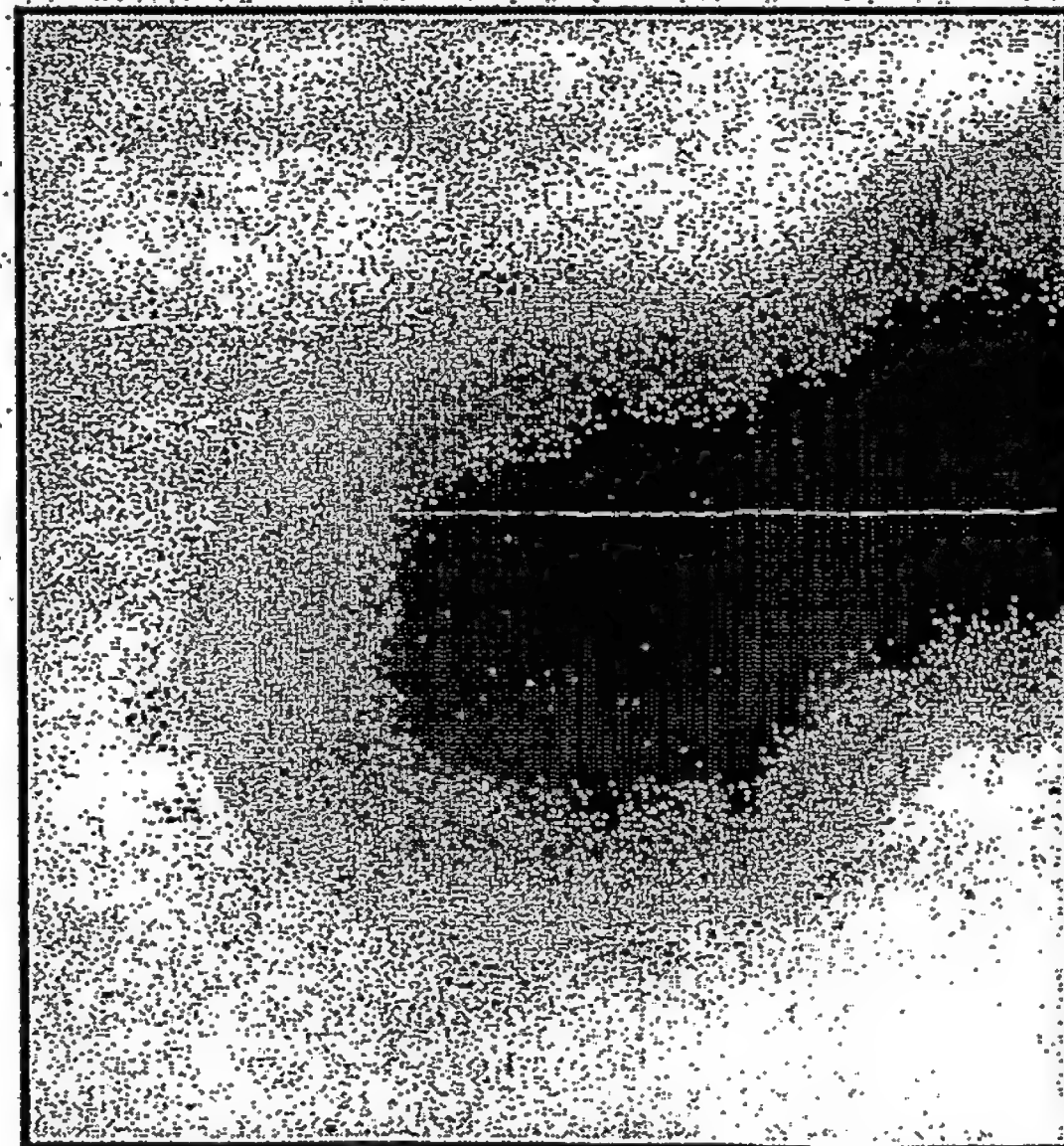
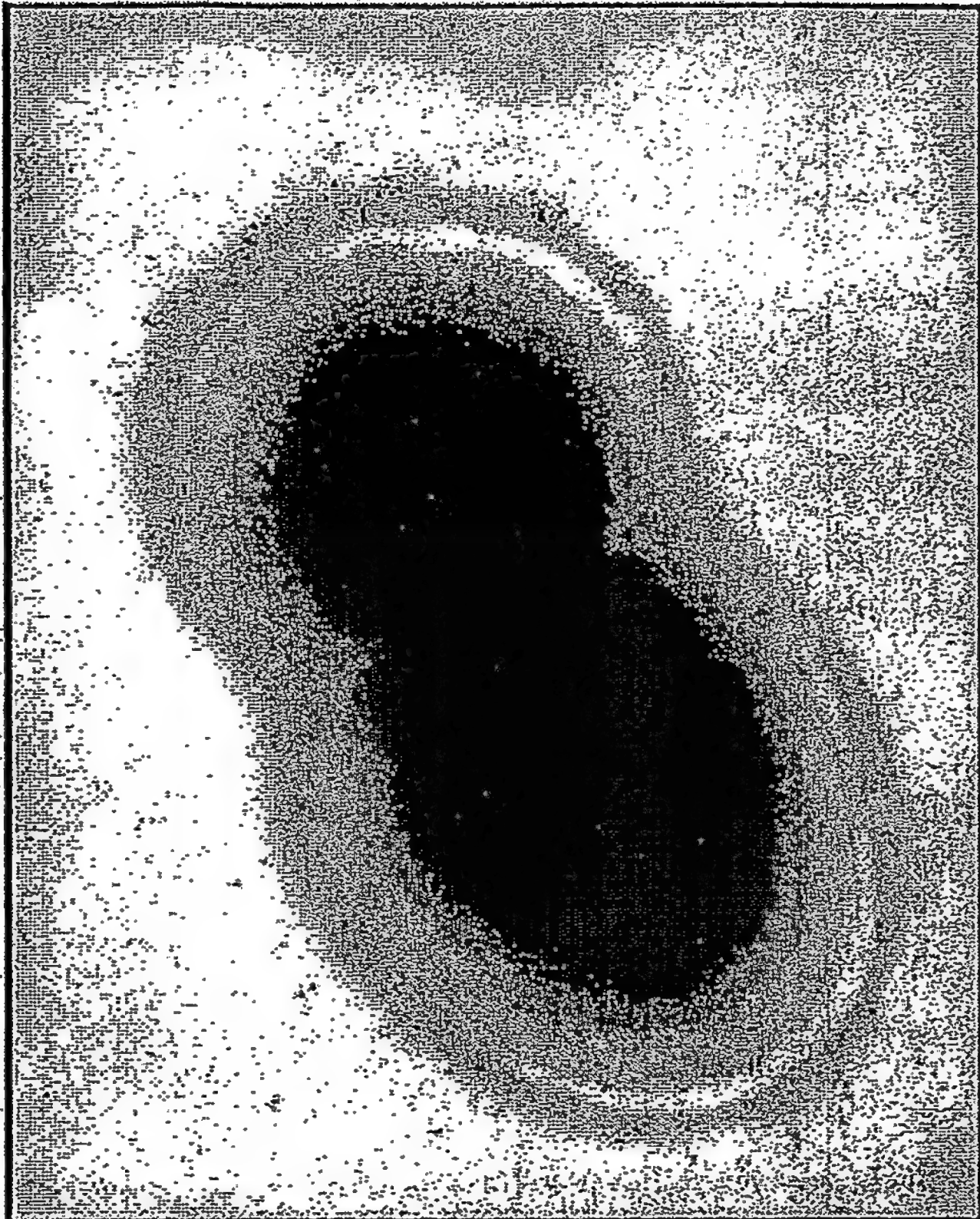
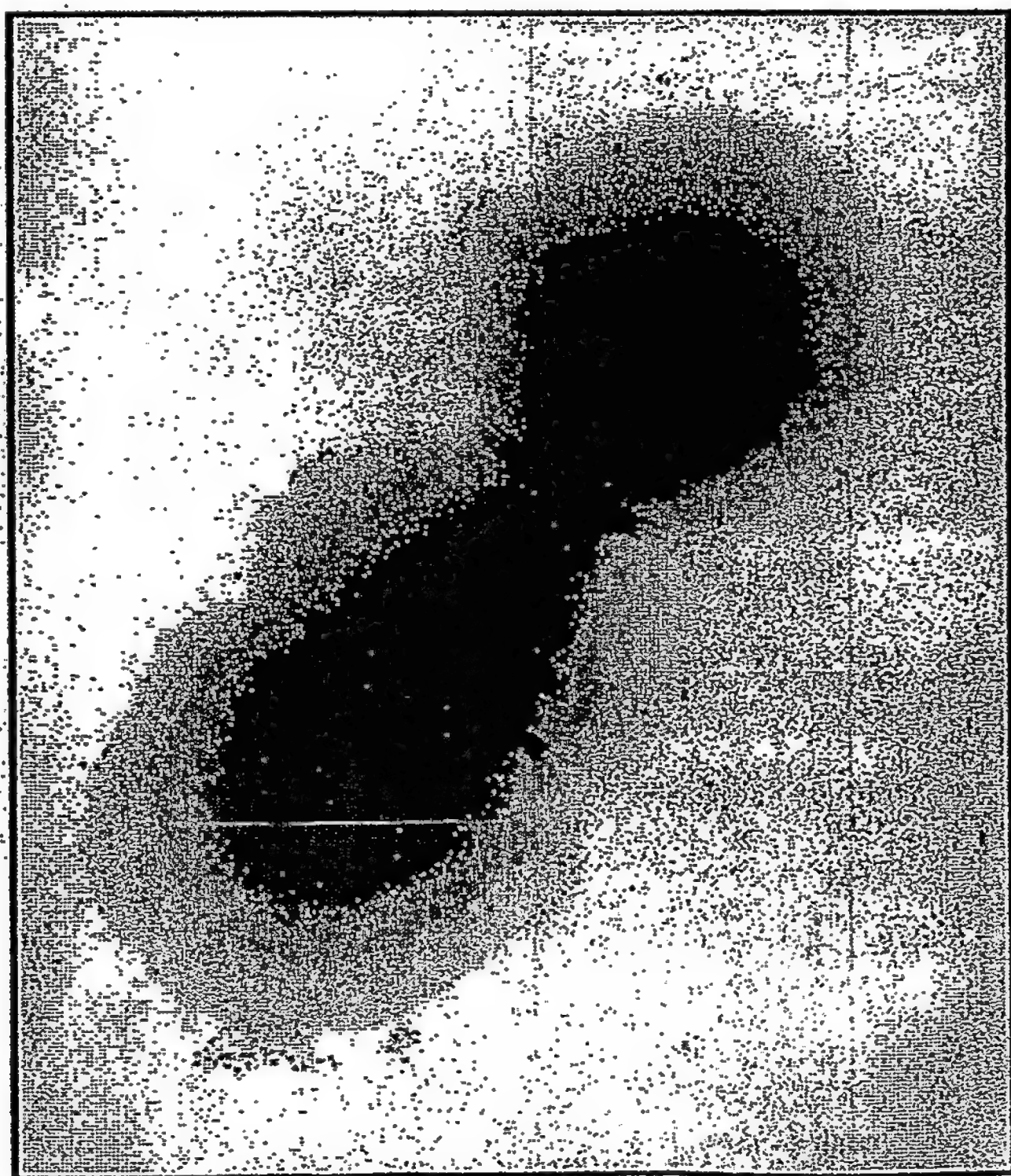
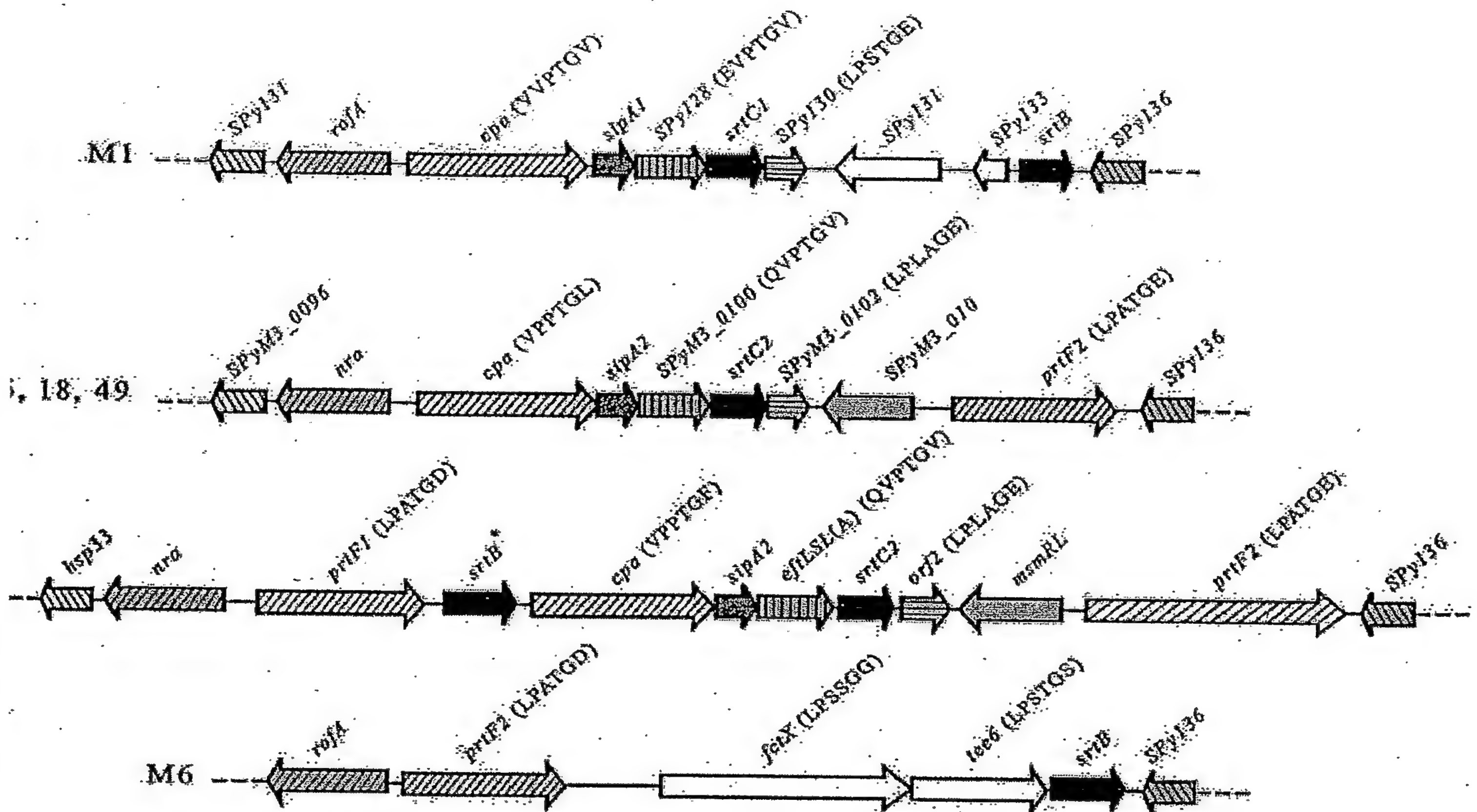


FIGURE 50





M1	1	MIVNLIKELDAIINVIYICFEEICLIPALGIVIALTYOGANATNYOQYK--KKGVOFDLLAENS DVERWI DVGTHII
M3	1	MYMTIVQVINKAIDTLILIFCLVVLFLAGFGLWDSYHLYOCADASNFKKFKTAQOQPKFEDLLALNEDVIGWLNIPGTHII
M5	1	YGMNIVQVINKAIDTLILIFCLVVLFLAGFGLWDSYHLYOCADASNFKKFKTAQOQPKFEDLLALNEDVIGWLNIPGTHII
M12	1	MYMTIVQVINKAIDTLILIFCLVVLFLAGFGLWDSYHLYOCADASNFKKFKTAQOQPKFEDLLALNEDVIGWLNIPGTHII
M18	1	MYMTIVQVINKAIDTLILIFCLVVLFLAGFGLWDSYHLYOCADASNFKKFKTAQOQPKFEDLLALNEDVIGWLNIPGTHII
M49	1	MYMTIVQVINKAIDTLILIFCLVVLFLAGFGLWDSYHLYOCADASNFKKFKTAQOQPKFEDLLALNEDVIGWLNIPGTHII
M1	80	YPLVOCENNLEYINKAVDGSVAMSGSIFLITRNHNDFTDDYSLIYCHHMAQNAFGEIPKFLKKDFFSKKNKATITETKFK
M3	82	YPLVOCENNLEYINKAVDGSVAMSGSIFLITRNHNDFTDDYSLIYCHHMAQNAFGEIPKFLKKDFFSKKNKATITETKFK
M5	82	YPLVOCENNLEYINKAVDGSVAMSGSIFLITRNHNDFTDDYSLIYCHHMAQNAFGEIPKFLKKDFFSKKNKATITETKFK
M12	82	YPLVOCENNLEYINKAVDGSVAMSGSIFLITRNHNDFTDDYSLIYCHHMAQNAFGEIPKFLKKDFFSKKNKATITETKFK
M18	82	YPLVOCENNLEYINKAVDGSVAMSGSIFLITRNHNDFTDDYSLIYCHHMAQNAFGEIPKFLKKDFFSKKNKATITETKFK
M49	82	YPLVOCENNLEYINKAVDGSVAMSGSIFLITRNHNDFTDDYSLIYCHHMAQNAFGEIPKFLKKDFFSKKNKATITETKFK
M1	161	KLKINIFACIKTDAFISLAFNFIIPVDLSSKHEEINHIKQSVYMRDILTTNESKFEVALSTCEENFTTICRTIVTFCIE
M3	163	KLTVTIFACLKTDAPDQLVFNPNATNQDQQLVLYISKRSKQKPKVKLKHHTKFAVAFSTCENFSTDNKRVIVVGTIOE
M5	163	KLTVTIFACLKTDAPDQLVFNPNATNQDQQLVLYISKRSKQKPKVKLKHHTKFAVAFSTCENFSTDNKRVIVVGTIOE
M12	163	KLTVTIFACLKTDAPDQLVFNPNATNQDQQLVLYISKRSKQKPKVKLKHHTKFAVAFSTCENFSTDNKRVIVVGTIOE
M18	163	KLTVTIFACLKTDAPDQLVFNPNATNQDQQLVLYISKRSKQKPKVKLKHHTKFAVAFSTCENFSTDNKRVIVVGTIOE
M49	163	KLTVTIFACLKTDAPDQLVFNPNATNQDQQLVLYISKRSKQKPKVKLKHHTKFAVAFSTCENFSTDNKRVIVVGTIOE

FIGURE 51



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GI-19224135	1	MNNKKLQKQDAPR-VSNRP	-----KQLTYTLGVFLMFLTLSSMRGAQSTIEGEEK
ORF78	1	-----LQKRDKTNYGSANNKR	-----ROTTIGLLKVFLTFVALIG-----IVG-----
GI-21909634	1	-----MOKRDKTNYGSANNKR	-----ROTTIGLLKVFLTFVALIGIVGFSIRAFG-----
GI-28810257	1	-----MOKRDKTNYGSANNKR	-----ROTTIGLLKVFLTFVALIGIVGFSIRAFG-----
GI-19745301	1	-----MOKRDKTNYGSANNKR	-----ROTTIGLLKVFLTFVALIGIVGFSIRAFG-----
GAS15	1	LRGEKMKKTRFPNKLNTLTORVLSKNS	KRFVTLGVFLMIFALVTSVVGAKTVEG-----
GI-19224135	53	RIEEVSVPKIKSPDD--AYPWYGYDSYDSSHEFYERFVAHDLRVNLNGSKSYQVYCFNL	
ORF78	39	-----	-----
GI-21909634	46	-AEEQSVPNKQSSVQ--DYPWYGYDSYSKGYEDYSPLNTYHNLKVNLDGSKEYQAYCFNL	
GI-28810257	46	-AEEQSVPNKQSSVQ--DYPWYGYDSYSKGYEDYSPLNTYHNLKVNLDGSKEYQAYCFNL	
GI-19745301	46	-AEEQST-----	-----
GAS15	58	-LVESSITPNAINPDSSSEYRWYGYESVVRGHEFYKQFRVAHDLRVNLEGSRSYQVYCFNL	
GI-19224135	111	NSHMPNEKNKAFSKQIENRVGDTGEVETNYAQTPKIRGESLNNKELSLIMYNAYPKNANGYM	
ORF78	39	-----	-----
GI-21909634	103	TKHFPSKSDSVRSQWYKKECTNENFIKLADKPRIEDGQLOONILRLIYNGYPNDRNGIM	
GI-28810257	103	TKHFPSKSDSVRSQWYKKECTNENFIKLADKPRIEDGQLOONILRLIYNGYPNDRNGIM	
GI-19745301	52	-----	-----
GAS15	117	KNAFFLGSDSVKKWYKKECTNENFIKLADKPRIEDGQLOONILRLIYNGYPNDRNGIM	
GI-19224135	171	DKLEPLNAILVTQQAIVYSDSSYGN-IKTLWASELKDGKIDFEQVKLMREAYSRLISDD	
ORF78	39	-----FSIRAE-----	-----
GI-21909634	163	KGIDPLNAILVTQNAIVYTDSSYISDTSKAFQOEETDLNLDSSQOLQLMRNALKRLINPK	
GI-28810257	163	KGIDPLNAILVTQNAIVYTDSSYISDTSKAFQOEETDLNLDSSQOLQLMRNALKRLINPK	
GI-19745301	52	-----	-----
GAS15	177	EGLEPLNAIRVTQQAIVYSDNAPISNPDESFKRESESNLVSTSQLSLMRQALNOLIDEN	
GI-19224135	230	LEETSKNKLEQGSKLNEVPODNS-----VONLLSAEYVPEPPAPGQSPPEPPVOTKNT	
ORF78	46	-----	-----AEEKSTETKNT
GI-21909634	223	EVESLPNOVPANVOLSIFQSSDKT-----FONLLSAEYVDPDTPPKPG-----EPPAKTENT	
GI-28810257	223	EVESLPNOVPANVOLSIFQSSDKT-----FONLLSAEYVDPDTPPKPG-----EPPAKTENT	
GI-19745301	52	-----	-----ETKNT
GAS15	237	LATKMPKQVPDDEQOLSIFESDNGDKYNKGXONLLSGGVPTKPPTPGDPPMPPNQPOTT	
GI-19224135	284	SVIIRKYAEGDYSKLLEGATLRLIGEDILDFQEKVFO-SNGTGERIELSNGTYTLTETSSP	
ORF78	57	SVIIRKYAEGDYSKLLEGATLRLIGEDILDFQEKVFO-SNGTGERIELSNGTYTLTETSSP	
GI-21909634	275	SVIIRKYAEGDYSKLLEGATLRLIAGIEGSGFOEKIFDSNKSCEKVELIAGTYVLSEIKPP	
GI-28810257	275	SVIIRKYAEGDYSKLLEGATLRLIAGIEGSGFOEKIFDSNKSCEKVELIAGTYVLSEIKPP	
GI-19745301	57	SVIIRKYAEGDYSKLLEGATLRLIAGIEGSGFOEKIFDSNKSCEKVELIAGTYVLSEIKPP	
GAS15	297	SVIIRKYAEGDYSKLLEGATLRLIGEDILDFQEKVFO-SNGTGERIELSNGTYTLTETSSP	
GI-19224135	344	DGYKIAEPIKFRVVKVFIQNDGSOVENPNKEVAEPYSVEAYSDMODSNYINPETETP	
ORF78	117	DGYKIAEPIKFRVVKVFIQNDGSOVENPNKEVAEPYSVEAYSDMODSNYINPETETP	
GI-21909634	335	DGYKIAEPIKFRVVKVFIQNDGSOVENPNKEVAEPYSVEAYSDMODSNYINPETETP	
GI-28810257	335	DGYKIAEPIKFRVVKVFIQNDGSOVENPNKEVAEPYSVEAYSDMODSNYINPETETP	
GI-19745301	117	DGYKIAEPIKFRVVKVFIQNDGSOVENPNKEVAEPYSVEAYSDMODSNYINPETETP	
GAS15	357	AGYSIAEPIKFRVVKVFIQNDGSOVENPNKEVAEPYSVEAYSDMODSNYINPETETP	
GI-19224135	404	YGRFYAKNKDKSSQVYCFNADLHSPPESEDGCGTIDPDISTMKVRYTHIAGSDLFKY	
ORF78	174	YGRFYAKNKDYDSSQIVYCFNANLKSPPDSEDHGATINPDFTTG-DIRYSHIAGSDLFKY	
GI-21909634	393	YGRFYAKNKDNGTQVYCFNADLHSPPESEDGCGTIDPDISTMKVRYTHIAGSDLFKY	
GI-28810257	393	YGRFYAKNKDNGTQVYCFNADLHSPPESEDGCGTIDPDISTMKVRYTHIAGSDLFKY	
GI-19745301	177	YGRFYAKNKANGTSQVYCFNADLHSPPESEDGCGTIDPDISTMKVRYTHIAGSDLFKY	
GAS15	413	YGRFYAKNKNGSSQVYCFNADLHSPPESEDGCGTIDPDISTMKVRYTHIAGSDLFKY	

FIGURE 52



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GI-19224135 464 ALRPRDTNPEDFLKHINKVIEKGYNKRGD--SYNGLTETQFRAATOLAIYYFTDSTDLKT  
ORF78 233 ANTARDEDEPOLFLKHVKNVIEKGYHKGQAIPYNSLTEAQFRAATOLAIYYFTDSVLDL--  
GI-21909634 453 AATPRDKDADFFLKHILKILDKGYKKRGD--TYKTLTEAQFRAATOLAIYYFTDSADLIT  
GI-28810257 453 AATPRDKDADFFLKHILKILDKGYKKRGD--TYKTLTERAQFRAATOLAIYYFTDSADLIT  
GI-19745301 237 ANNPRASTNDELLSCVNVLEKGYRDDST--TYANLTSVEFRAATOLAIYYFTDSVLDLND  
GAS15 472 TVKPRDEDEPDTFLKHINKVIEKGYREKQAIBYSCLTETQLRAATOLAIYYFTDSAEID-

GI-19224135 522 LKTYNNGKGYHGFESMDEKTLAVIKELINYAOD-NSAPQLTNLDFFVPNNISKYOSLIGTE  
ORF78 291 --TKDRIKDEHGFGDMNDOTLGVAKKIVEYALS-DEDSKLTNLDFFVPNNISKYOSLIGTE  
GI-21909634 511 LKTYNDNKGYPHGFDEKLDDEATLAVVHELTIAED-VLEFMTQNLDFVPNNSSRYOALIGTO  
GI-28810257 511 LKTYNDNKGYPHGFDEKLDDEATLAVVHELTIAED-VLEFMTQNLDFVPNNSSRYOALIGTO  
GI-19745301 295 LADY-----HGFGALTTEALNATLEIVAYAEADRANLPNISNLDFVPNNISKYOSLIGTO  
GAS15 531 ---KDKLNDYHGFGDMNDSTLAVANILMEYAOD-SNPQLTDLDFVPNNISKYOSLIGTO

GI-19224135 581 YHPDDLVDVIRMEDNKQEVIPVTHSLTVKRTVVGELGDMINGFQFELBLNDKTCOPIVNT  
ORF78 348 YHPDDLVDVIRMEDNKQEVIPVTHSLTVKRTVVGELGDMINGFQFELBLNDKTCOPIVNT  
GI-21909634 570 YHPNELIDVISMEDNKQPIIPITHKLTISKVTGTIADNKKEFNFEIHLKSSDGOAISGT  
GI-28810257 570 YHPNELIDVISMEDNKQPIIPITHKLTISKVTGTIADNKKEFNFEIHLKSSDGOAISGT  
GI-19745301 349 YHPESLVDIIRMEDNKQPIIPITHKLTISKVTGTIADNKKEFNFEIHLKSSDGOAISGT  
GAS15 587 YHPEDLVDIIRMEDNK-EVIPVTHNLTIRKVTGCLAGDFTNDFHFEIHLNNKQELLSQT

GI-19224135 641 LKTNNOBLVAKDCKYSENLNKGDTIRIEGLPTGYSYTEKETBAKDYIVTVDNVVSQEAQS  
ORF78 408 LKTNNOBLVAKDCKYSENLNKGDTIRIEGLPTGYSYTEKETBAKDYIVTVDNVVSQEAQS  
GI-21909634 630 YPTNSGELTVIDGKATFLLNDGESLIVEGLPSGYSYEITETGASDYEVSNGKNAPDGKA  
GI-28810257 630 YPTNSGELTVIDGKATFLLNDGESLIVEGLPSGYSYEITETGASDYEVSNGKNAPDGKA  
GI-19745301 409 YPTNSGELTVIDGKATFLLNDGESLIVEGLPSGYSYEITETGASDYEVSNGKNAPDGKA  
GAS15 646 VKTDKTNLEFKDCKATINLNKGESLTLOGLPFGYSYLVKETDSEGYKVKVNSQEVANATV

GI-19224135 701 ASENVTACKEVTFENRRDLVPPTGCLTDDGGTYLWLLLLLVFFGLLWVLFGRKGLND-  
ORF78 468 ASENVTACKEVTFENRRDLVPPTGCLTDDGATYLVWLLLLLVFFGLLWVLFGRKGLND-  
GI-21909634 690 TRASVKEDETVAFENRRDLVPPTGCLTDDGATYLVWLLLLLVFFGLLWVLFGRKGLTKK--  
GI-28810257 690 TRASVKEDETVAFENRRDLVPPTGCLTDDGATYLVWLLLLLVFFGLLWVLFGRKGLTKK--  
GI-19745301 469 TRASVKEDETVAFENRRDLVPPTGCLTDDGATYLVWLLLLLVFFGLLWVLFGRKGLND-  
GAS15 706 SKTGITSDETEAFENRRDLVPPTGCLTDDGATYLVWLLLLLVFFGLLWVLFGRKGLTKK--

FIGURE 52A



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GI-50913503 1 MVSSYMFARCEKMNKMLNKEAGFLVHTKRRKRFVTLVGVFFLLACAGATGFGQVAY

GI-19224134 61 AADERTVFNFKSPDPDYPIYGYDSY-----RGIFARYHNLKVNKGSRYQAYCFNITK  
GI-50913503 61 AADERTVPSHS SPNPEFPWYGYDAYGREYPCYNIMTRYHDLRVNLNGSRSYQVYCFNITQS

GI-19224134 115 YEPFRPTYSTTNMFKKIDGSCSAFKSYAANPRVLDENLCKLEKNILNVTYNGYKSNANGE  
GI-50913503 121 NYPSSQKNSFIKNNFKKIEGCKSFVDYAHTTKLGKE---ELEQRRLSLEYNEYPNDANGY

GI-19224134 175 MNGTIEDLNAILVTONATHYYSDSAPLNDVNKMIEREVRNGEISESQVTLMREALKKLIDP  
GI-50913503 178 MKGLEHLNAILVTQYAVHYSDNS-QYQFETLIESEANECKISR SQVTLMREALKKLIDP

GI-19224134 235 NLEATAANKIPSGYRLNIFKSENEAYQNLLSAEYVPDDPPKPGDTSEHNPKTPELDGTPI  
GI-50913503 237 NLEATAVNKIPSGYRLNIFESENEAYQNLLSAEYVPDDPPKPGGETSEHNPKTPELDGTPI

GI-19224134 295 PEDPKRPDESSEPALPPLMPELDGEVPEVPSESLEPALPPLMPELDGEVPEVPSESLE  
GI-50913503 297 PEDPKRPDDNLEPTLPFVM-----

GI-19224134 355 PALPPLMPELDGEVPEVPSESLEPALPPLMPELDGEVPEVPSESLEPALPPLMPELDG  
GI-50913503 316 -----LDGEVPEVPSESLEPALPPLMPELDG

GI-19224134 415 BEVPEKPSVDLPFIEVPRYEFNNKDOSPLACESGETEYITEVYCNQONPVDIDKILPNETG  
GI-50913503 343 QEVPEKPSIDLPIEVPRYEFNNKDOSPLACESGETEYITEVYCNQONPVDIDKILPNETG

GI-19224134 475 FSGNMVETEDTKEPEVLMGGQSESVEFTKDTQTGMSGOTTPOVETEDTKEPEVLMGGQSE  
GI-50913503 403 FSGNMVETEDTKEPEVLMGGQSESVEFTKDTQTGMSGOTTPOVETEDTKEPEVLMGGQSE

GI-19224134 535 SVEFTKDTQTGMSGOTTPOVETEDTKEPEVLMGGQSESVEFTKDTQTGMSGOTTPOVETE  
GI-50913503 463 SVEFTKDTQTGMSGOTTPOIETEDTKEPEVLMGGQSESVEFTKDTQTGMSGOTTPOIETE

GI-19224134 595 DTKEPEVLMGGQSESVEFTKDTQTGMSGFSETATIVEDTRPKLVFHFDMNEPKVEENREK  
GI-50913503 523 DTKEPEVLMGGQSESVEFTKDTQTGMSGFSETATIVEDTRPKLVFHFDMNEPKVEENREK

GI-19224134 655 PTKNITPILPATGDIENVLAFLGILILSVLSIFSLLNKKNKVV-  
GI-50913503 583 PTKNITPILPATGDIENVLAFLGILILSVLSIFSLLNKKNKSNKVV

FIGURE 53



GI-19745307	1	MTOKNSY	FLLSLTGFILGLLLVFIGLSGVSVGHAETRNGANKQG	
ORF84		WO 2006/078318	NSYNLSFLLSLTGFILGLLLVFIGLSGVSVGHAETRNGANKQGAPCT/US2005/027239	
GI-28810263	1	MTOKNSYKLSFLLSLTGFILGLLLVFIGLSGVSVGHAETRNGANKQGAPEIKNN		
GI-21909640	1			
GI-19224141	1	MTOKNSYKLSFLLSLTGFILGLLLVFIGLSGVSVGHAETRNGANKQGSFEIKKVDQNNKP		
GI-19745307	55			KSQEEYNYE
ORF84	55			KSQEEYNYE
GI-28810263	55			KSQEEYNYE
GI-21909640	1			
GI-19224141	61	LPGATFSLTSKDGKGTSVQTFSTNDKGIVDAQNLQPGTYTLKEETAPDGYD		KSRTWTVT
GI-19745307	64	VYDN		RNI
ORF84	64	VYDN		RNI
GI-28810263	64	VYDN		RNI
GI-21909640	1			
GI-19224141	121	VYEN	GYTKLVENPYNGEIIISKAGSKDVSSSLQLENPKMSVVSKYGKTEVSSGAADFY	RNI
GI-19745307	71	LQDGEHKLEIKRVDGTGKTYQC	FCFQLTNNFF	TAQGVSKKLYKKLSS
ORF84	71	LQDGEHKLEIKRVDGTGKTYQC	FCFQLTNNFF	TAQGVSKKLYKKLSS
GI-28810263	71	LQDGEHKLEIKRVDGTGKTYQC	FCFQLTNNFF	TAQGVSKKLYKKLSS
GI-21909640	1			MSS
GI-19224141	181	AAYFKMSF	ENKQDKSETINP	CDT
			EVLDRLNPKGISQDEPIFYDSANS	PLAIGKYH
GI-19745307	118			SDEETLK
ORF84	118			SDEETLK
GI-28810263	118			SDEETLK
GI-21909640	4			SDEETLK
GI-19224141	241	AENHQLIYTFTDYIAGLDKVQLSAELSLFLENKE		EVLENTSISNFKSTIGGQETIKGTVN
GI-19745307	125			QYASKYTSNRRGDTSC
ORF84	125			QYASKYTSNRRGDTSC
GI-28810263	125			QYASKYTSNRRGDTSC
GI-21909640	11			QYASKYTSNRRGDTSC
GI-19224141	301	VLYGNESTKES	NYITNGLSNVGG	SIESYNTETGEFVWYVYNPNRTNIPYATMNLWGFR
GI-19745307	141			NLRKQIAKVLTEGYPT
ORF84	141			NLRKQIAKVLTEGYPT
GI-28810263	141			NLRKQIAKVLTEGYPT
GI-21909640	27			NLRKQIAKVLTEGYPT
GI-19224141	361	ARSNTSDLENDANTSSAELGEIQVYEVPEGEKLPSSYGVDVTK		LEDDIACLGNGFOM
GI-19745307	157	NKS	DWLN	GLTENEKIEVTQDAIWYF
ORF84	157	NKS	DWLN	GLTENEKIEVTQDAIWYF
GI-28810263	157	NKS	DWLN	GLTENEKIEVTQDAIWYF
GI-21909640	43	NKS	DWLN	GLTENEKIEVTQDAIWYF
GI-19224141	421	TRQRIDE	GNNIQNKAFIIKVTGKTDQSGKPLVVQSNLASFR	CASEYAAFTPVGGNVYEQ
GI-19745307	182	TETTVPADR		SYTNRRNVNSQKMKEVYQKLI
ORF84	182	TETTVPADR		SYTNRRNVNSQKMKEVYQKLI
GI-28810263	182	TETTVPADR		SYTNRRNVNSQKMKEVYQKLI
GI-21909640	68	TETTVPADR		SYTNRRNVNSQKMKEVYQKLI
GI-19224141	481	NETALSPSK	GSGSGKSEFTKPSITVANLRVAQLRFKKMST	DNVPLPEAAFEELRSSNGNS
GI-19745307	233			LQAVISVEPVIESLPITS
ORF84	233			LQAVISVEPVIESLPITS
GI-28810263	233			LQAVISVEPVIESLPITS
GI-21909640	119			LQAVISVEPVIESLPITS
GI-19224141	541	QKLEASSNT	QGEVHFKDLTS	GTYDLYETKAPKGYQOVTEKEATVTVDTTNPAEMVTWGS

FIGURE 54